## SEQUENCE LISTING

- <110> Cellzome AG
- <120> Protein complexes of the TIP60 transcriptional activator protein as well as components, fragments and derivatives thereof and methods for using the same
- <130> Protein complexes of the TIP60 transcriptional activator protein as well as components, fragments and derivatives thereof and methods for using the same
- <150> EP02016110.5 <151> 2002-07-19
- <150> EP03101321.2
- <151> 2003-05-12
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- <170> PatentIn version 3.1
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- Val Ile Gln Asn Pro Gly Pro Arg His Pro Glu Ala Ala Ser Ala Ala
- Pro Pro Gly Ala Ser Leu Leu Leu Gln Gln Gln Gln Gln Gln Gln 55
- 75 65 70
- Ser Pro Arg Gln Gln Gln Gln Gln Gln Gly Glu Asp Gly Ser Pro Gln
- Ala His Arg Arg Gly Pro Thr Gly Tyr Leu Val Leu Asp Glu Glu Gln
- Gln Pro Ser Gln Pro Gln Ser Ala Leu Glu Cys His Pro Glu Arg Gly
- Cys Val Pro Glu Pro Gly Ala Ala Val Ala Ala Ser Lys Gly Leu Pro 130 135
- Gln Gln Leu Pro Ala Pro Pro Asp Glu Asp Asp Ser Ala Ala Pro Ser
- Thr Leu Ser Leu Leu Gly Pro Thr Phe Pro Gly Leu Ser Ser Cys Ser 165

Ala Asp Leu Lys Asp Ile Leu Ser Glu Ala Ser Thr Met Gln Leu Leu 180 185 190

Gln Gln Gln Gln Glu Ala Val Ser Glu Gly Ser Ser Ser Gly Arg 195 200 205

Ala Arg Glu Ala Ser Gly Ala Pro Thr Ser Ser Lys Asp Asn Tyr Leu 210 215 220

Gly Gly Thr Ser Thr Ile Ser Asp Asn Ala Lys Glu Leu Cys Lys Ala 225 230 235 240

Val Ser Val Ser Met Gly Leu Gly Val Glu Ala Leu Glu His Leu Ser 245 250 255

Pro Gly Glu Gln Leu Arg Gly Asp Cys Met Tyr Ala Pro Leu Leu Gly 260 265 270

Val Pro Pro Ala Val Arg Pro Thr Pro Cys Ala Pro Leu Ala Glu Cys 275 280 285

Lys Gly Ser Leu Leu Asp Asp Ser Ala Gly Lys Ser Thr Glu Asp Thr 290 295 300

Ala Glu Tyr Ser Pro Phe Lys Gly Gly Tyr Thr Lys Gly Leu Glu Gly 305 310 315 320

Glu Ser Leu Gly Cys Ser Gly Ser Ala Ala Gly Ser Ser Gly Thr 325 330 335

Leu Glu Leu Pro Ser Thr Leu Ser Leu Tyr Lys Ser Gly Ala Leu Asp 340 345

Glu Ala Ala Tyr Gln Ser Arg Asp Tyr Tyr Asn Phe Pro Leu Ala 355 360 365

Leu Ala Gly Pro Pro Pro Pro Pro Pro Pro Pro His Pro His Ala Arg 370 375 380

Ile Lys Leu Glu Asn Pro Leu Asp Tyr Gly Ser Ala Trp Ala Ala Ala
385 390 395 400

Ala Ala Gln Cys Arg Tyr Gly Asp Leu Ala Ser Leu His Gly Ala Gly 405 410 415

Ala Ala Gly Pro Gly Ser Gly Ser Pro Ser Ala Ala Ala Ser Ser Ser 420 425 430

Trp His Thr Leu Phe Thr Ala Glu Glu Gly Gln Leu Tyr Gly Pro Cys 435 440 445

Gly Gly Gly Gly Gly Gly Glu Ala Gly Ala Val Ala Pro Tyr 465 470 475 480

Gly Tyr Thr Arg Pro Pro Gln Gly Leu Ala Gly Gln Glu Ser Asp Phe
485 490 495

Thr Ala Pro Asp Val Trp Tyr Pro Gly Gly Met Val Ser Arg Val Pro 500 505 510

Tyr Pro Ser Pro Thr Cys Val Lys Ser Glu Met Gly Pro Trp Met Asp 515 520 525

Ser Tyr Ser Gly Pro Tyr Gly Asp Met Arg Leu Glu Thr Ala Arg Asp 530 535 540

His Val Leu Pro Ile Asp Tyr Tyr Phe Pro Pro Gln Lys Thr Cys Leu 545 550 555 560

Ile Cys Gly Asp Glu Ala Ser Gly Cys His Tyr Gly Ala Leu Thr Cys 565 570 575

Gly Ser Cys Lys Val Phe Phe Lys Arg Ala Ala Glu Gly Lys Gln Lys 580 585 590

Tyr Leu Cys Ala Ser Arg Asn Asp Cys Thr Ile Asp Lys Phe Arg Arg 595 600 605

Lys Asn Cys Pro Ser Cys Arg Leu Arg Lys Cys Tyr Glu Ala Gly Met 610 620

Thr Leu Gly Ala Arg Lys Leu Lys Lys Leu Gly Asn Leu Lys Leu Gln 625 630 635 640

Glu Glu Gly Glu Ala Ser Ser Thr Thr Ser Pro Thr Glu Glu Thr Thr 645 650 655

Gln Lys Leu Thr Val Ser His Ile Glu Gly Tyr Glu Cys Gln Pro Ile 660 665 670

Phe Leu Asn Val Leu Glu Ala Ile Glu Pro Gly Val Val Cys Ala Gly 675 680 685

His Asp Asn Asn Gln Pro Asp Ser Phe Ala Ala Leu Leu Ser Ser Leu 690 695 700

Asn Glu Leu Gly Glu Arg Gln Leu Val His Val Val Lys Trp Ala Lys 705 710 715 720

Ala Leu Pro Gly Phe Arg Asn Leu His Val Asp Asp Gln Met Ala Val 730 Ile Gln Tyr Ser Trp Met Gly Leu Met Val Phe Ala Met Gly Trp Arg Ser Phe Thr Asn Val Asn Ser Arg Met Leu Tyr Phe Ala Pro Asp Leu Val Phe Asn Glu Tyr Arg Met His Lys Ser Arg Met Tyr Ser Gln Cys Val Arg Met Arg His Leu Ser Gln Glu Phe Gly Trp Leu Gln Ile Thr Pro Gln Glu Phe Leu Cys Met Lys Ala Leu Leu Phe Ser Ile Ile Pro Val Asp Gly Leu Lys Asn Gln Lys Phe Phe Asp Glu Leu Arg Met Asn Tyr Ile Lys Glu Leu Asp Arg Ile Ile Ala Cys Lys Arg Lys Asn 835 840 845 Pro Thr Ser Cys Ser Arg Arg Phe Tyr Gln Leu Thr Lys Leu Leu Asp 855 Ser Val Gln Pro Ile Ala Arg Glu Leu His Gln Phe Thr Phe Asp Leu 870 Leu Ile Lys Ser His Met Val Ser Val Asp Phe Pro Glu Met Met Ala 885 890 Glu Ile Ile Ser Val Gln Val Pro Lys Ile Leu Ser Gly Lys Val Lys 905 Pro Ile Tyr Phe His Thr Gln <210> 2 <211> 375 <212> PRT <213> Homo sapiens <400> 2 Met Asp Asp Asp Ile Ala Ala Leu Val Val Asp Asn Gly Ser Gly Met Cys Lys Ala Gly Phe Ala Gly Asp Asp Ala Pro Arg Ala Val Phe Pro Ser Ile Val Gly Arg Pro Arg His Gln Gly Val Met Val Gly Met Gly

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Leu Thr Leu Lys Tyr Pro Ile Glu His Gly Ile Val Thr Asn Trp Asp 65 70 75 80

Asp Met Glu Lys Ile Trp His His Thr Phe Tyr Asn Glu Leu Arg Val 85 90 95

Ala Pro Glu Glu His Pro Val Leu Leu Thr Glu Ala Pro Leu Asn Pro 100 105 110

Lys Ala Asn Arg Glu Lys Met Thr Gln Ile Met Phe Glu Thr Phe Asn 115 120 125

Thr Pro Ala Met Tyr Val Ala Ile Gln Ala Val Leu Ser Leu Tyr Ala 130 135 140

Ser Gly Arg Thr Thr Gly Ile Val Met Asp Ser Gly Asp Gly Val Thr 145 150 155 160

His Thr Val Pro Ile Tyr Glu Gly Tyr Ala Leu Pro His Ala Ile Leu 165 170 175

Arg Leu Asp Leu Ala Gly Arg Asp Leu Thr Asp Tyr Leu Met Lys Ile 180 185 190

Leu Thr Glu Arg Gly Tyr Ser Phe Thr Thr Ala Glu Arg Glu Ile 195 200 205

Val Arg Asp Ile Lys Glu Lys Leu Cys Tyr Val Ala Leu Asp Phe Glu 210 215 220

Gln Glu Met Ala Thr Ala Ala Ser Ser Ser Ser Leu Glu Lys Ser Tyr 225 230 235 240

Glu Leu Pro Asp Gly Gln Val Ile Thr Ile Gly Asn Glu Arg Phe Arg 245 250 255

Cys Pro Glu Ala Leu Phe Gln Pro Ser Phe Leu Gly Met Glu Ser Cys 260 265 270

Gly Ile His Glu Thr Thr Phe Asn Ser Ile Met Lys Cys Asp Val Asp 275 280 285

Ile Arg Lys Asp Leu Tyr Ala Asn Thr Val Leu Ser Gly Gly Thr Thr 290 295 300

Met Tyr Pro Gly Ile Ala Asp Arg Met Gln Lys Glu Ile Thr Ala Leu 305 310 315 320

Ala Pro Ser Thr Met Lys Ile Lys Ile Ile Ala Pro Pro Glu Arg Lys

Tyr Ser Val Trp Ile Gly Gly Ser Ile Leu Ala Ser Leu Ser Thr Phe

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Ile Val His Arg Lys Cys Phe 370

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Pro Lys Val Asp Phe Pro Thr Ala Ile Gly Met Val Val Glu Arg Asp

Asp Gly Ser Thr Leu Met Glu Ile Asp Gly Asp Lys Gly Lys Gln Gly

Gly Pro Thr Tyr Tyr Ile Asp Thr Asn Ala Leu Arg Val Pro Arg Glu

Asn Met Glu Ala Ile Ser Pro Leu Lys Asn Gly Met Val Glu Asp Trp

Asp Ser Phe Gln Ala Ile Leu Asp His Thr Tyr Lys Met His Val Lys

Ser Glu Ala Ser Leu His Pro Val Leu Met Ser Glu Ala Pro Trp Asn 120

Thr Arg Ala Lys Arg Glu Lys Leu Thr Glu Leu Met Phe Glu His Tyr 135

Asn Ile Pro Ala Phe Phe Leu Cys Lys Thr Ala Val Leu Thr Ala Phe 145

Ala Asn Gly Arg Ser Thr Gly Leu Ile Leu Asp Ser Gly Ala Thr His 175

Thr Thr Ala Ile Pro Val His Asp Gly Tyr Val Leu Gln Gln Gly Ile

180 185 190

Val Lys Ser Pro Leu Ala Gly Asp Phe Ile Thr Met Gln Cys Arg Glu 195 200 205

Leu Phe Gln Glu Met Asn Ile Glu Leu Val Pro Pro Tyr Met Ile Ala 210 215 220

Ser Lys Glu Ala Val Arg Glu Gly Ser Pro Ala Asn Trp Lys Arg Lys 225 230 235 240

Glu Lys Leu Pro Gln Val Thr Arg Ser Trp His Asn Tyr Met Cys Asn 245 250 255

Cys Val Ile Gln Asp Phe Gln Ala Ser Val Leu Gln Val Ser Asp Ser 260 265 270

Thr Tyr Asp Glu Gln Val Ala Ala Gln Met Pro Thr Val His Tyr Glu 275 280 285

Phe Pro Asn Gly Tyr Asn Cys Asp Phe Gly Ala Glu Arg Leu Lys Ile 290 295 300

Pro Glu Gly Leu Phe Asp Pro Ser Asn Val Lys Gly Leu Ser Gly Asn 305 310 315 320

Thr Met Leu Gly Val Ser His Val Val Thr Thr Ser Val Gly Met Cys 325 330 335

Asp Ile Asp Ile Arg Pro Gly Leu Tyr Gly Ser Val Ile Val Ala Gly 340 345 350

Gly Asn Thr Leu Ile Gln Ser Phe Thr Asp Arg Leu Asn Arg Glu Leu 355 360 365

Ser Gln Lys Thr Pro Pro Ser Met Arg Leu Lys Leu Ile Ala Asn Asn 370 380

Thr Thr Val Glu Arg Arg Phe Ser Ser Trp Ile Gly Gly Ser Ile Leu 385 390 395 400

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<210> 4

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Pro Glu Val Glu Val Cys Leu Phe His Ala Met Leu Gly His Lys Pro 35 40 45

Val Gly Val Asn Arg His Phe His Met Ile Cys Ile Arg Asp Lys Phe 50 55 60

Ser Gln Asn Ile Gly Arg Gln Val Pro Ser Lys Val Ile Trp Asp His 65 70 75 80

Leu Ser Thr Met Tyr Asp Met Gln Ala Leu His Glu Ser Glu Ile Leu 85 90 95

Pro Phe Pro Asn Pro Glu Arg Asn Phe Val Leu Pro Glu Glu Ile Ile 100 105 110

Gln Glu Val Arg Glu Gly Lys Val Met Ile Glu Glu Glu Met Lys Glu 115 120 125

Glu Met Lys Glu Asp Val Asp Pro His Asn Gly Ala Asp Asp Val Phe 130 135 140

Ser Ser Ser Gly Ser Leu Gly Lys Ala Ser Glu Lys Ser Ser Lys Asp 145 150 155 160

Lys Glu Lys Asn Ser Ser Asp Leu Gly Cys Lys Glu Gly Ala Asp Lys 165 170 175

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<210> 5

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Glu Gly Asp Ala Ala Ser Gly Thr Ile Ser Lys Lys Asp Ile Ile Asn 20 25 30

Pro Asp Lys Lys Ser Lys Ser Ser Glu Thr Leu Thr Phe Lys

Arg Pro Glu Gly Met His Arg Glu Val Tyr Ala Leu Leu Tyr Ser Asp Lys Lys Asp Ala Pro Pro Leu Leu Pro Ser Asp Thr Gly Gln Gly Tyr Arg Thr Val Lys Ala Lys Leu Gly Ser Lys Lys Val Arg Pro Trp Lys 90 Trp Met Pro Phe Thr Asn Pro Ala Arg Lys Asp Gly Ala Met Phe Phe 105 His Trp Arg Arg Ala Ala Glu Glu Gly Lys Asp Tyr Pro Phe Ala Arg Phe Asn Lys Thr Val Gln Val Pro Val Tyr Ser Glu Gln Glu Tyr Gln Leu Tyr Leu His Asp Asp Ala Trp Thr Lys Ala Glu Thr Asp His Leu 155 150 Phe Asp Leu Ser Arg Arg Phe Asp Leu Arg Phe Val Val Ile His Asp Arg Tyr Asp His Gln Gln Phe Lys Lys Arg Ser Val Glu Asp Leu Lys Glu Arg Tyr Tyr His Ile Cys Ala Lys Leu Ala Asn Val Arg Ala Val Pro Gly Thr Asp Leu Lys Ile Pro Val Phe Asp Ala Gly His Glu Arg 210 215 Arg Arg Lys Glu Gln Leu Glu Arg Leu Tyr Asn Arg Thr Pro Glu Gln 235 225 Val Ala Glu Glu Glu Tyr Leu Leu Gln Glu Leu Arg Lys Ile Glu Ala 250 245 Arg Lys Lys Glu Arg Glu Lys Arg Ser Gln Asp Leu Gln Lys Leu Ile Thr Ala Ala Asp Thr Thr Ala Glu Gln Arg Arg Thr Glu Arg Lys Ala 280 275 Pro Lys Lys Leu Pro Gln Lys Lys Glu Ala Glu Lys Pro Ala Val Pro Glu Thr Ala Gly Ile Lys Phe Pro Asp Phe Lys Ser Ala Gly Val

315

Thr Leu Arg Ser Gln Arg Met Lys Leu Pro Ser Ser Val Gly Gln Lys 325 330 335

Lys Ile Lys Ala Leu Glu Gln Met Leu Leu Glu Leu Gly Val Glu Leu 340 345 350

Ser Pro Thr Pro Thr Glu Glu Leu Val His Met Phe Asn Glu Leu Arg 355 360 365

Ser Asp Leu Val Leu Leu Tyr Glu Leu Lys Gln Ala Cys Ala Asn Cys 370 375 380

Glu Tyr Glu Leu Gln Met Leu Arg His Arg His Glu Ala Leu Ala Arg 385 390 395 400

Ala Gly Val Leu Gly Gly Pro Ala Thr Pro Ala Ser Gly Pro Gly Pro 405 410 415

Ala Ser Ala Glu Pro Ala Val Thr Glu Pro Gly Leu Gly Pro Asp Pro 420 425 430

Lys Asp Thr Ile Ile Asp Val Val Gly Ala Pro Leu Thr Pro Asn Ser 435 440 445

Arg Lys Arg Arg Glu Ser Ala Ser Ser Ser Ser Ser Val Lys Lys Ala 450 455 460

Lys Lys Pro 465

<210> 6

<211> 463

<212> PRT

<213> Homo sapiens

<400> 6

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Thr Arg Ile Glu Arg Ile Gly Ala His Ser His Ile Arg Gly Leu Gly 20 25 30

Leu Asp Asp Ala Leu Glu Pro Arg Gln Ala Ser Gln Gly Met Val Gly 35 40 45

Gln Leu Ala Arg Arg Ala Ala Gly Val Val Leu Glu Met Ile Arg  $50 \hspace{1cm} 55 \hspace{1cm} 60$ 

Glu Gly Lys Ile Ala Gly Arg Ala Val Leu Ile Ala Gly Gln Pro Gly 65 70 75 80

Thr Gly Lys Thr Ala Ile Ala Met Gly Met Ala Gln Ala Leu Gly Pro Asp Thr Pro Phe Thr Ala Ile Ala Gly Ser Glu Ile Phe Ser Leu Glu Met Ser Lys Thr Glu Ala Leu Thr Gln Ala Phe Arg Arg Ser Ile Gly Val Arg Ile Lys Glu Glu Thr Glu Ile Ile Glu Gly Glu Val Val Glu Ile Gln Ile Asp Arg Pro Ala Thr Gly Thr Gly Ser Lys Val Gly Lys Leu Thr Leu Lys Thr Thr Glu Met Glu Thr Ile Tyr Asp Leu Gly Thr Lys Met Ile Glu Ser Leu Thr Lys Asp Lys Val Gln Ala Gly Asp Val Ile Thr Ile Asp Lys Ala Thr Gly Lys Ile Ser Lys Leu Gly Arg Ser 195 Phe Thr Arg Ala Arg Asp Tyr Asp Ala Met Gly Ser Gln Thr Lys Phe 210 Val Gln Cys Pro Asp Gly Glu Leu Gln Lys Arg Lys Glu Val Wal His Thr Val Ser Leu His Glu Ile Asp Val Ile Asn Ser Arg Thr Gln Gly Phe Leu Ala Leu Phe Ser Gly Asp Thr Gly Glu Ile Lys Ser Glu Val Arg Glu Gln Ile Asn Ala Lys Val Ala Glu Trp Arg Glu Glu Gly Lys Ala Glu Ile Ile Pro Gly Val Leu Phe Ile Asp Glu Val His Met Leu 295 Asp Ile Glu Ser Phe Ser Phe Leu Asn Arg Ala Leu Glu Ser Asp Met 315 Ala Pro Val Leu Ile Met Ala Thr Asn Arg Gly Ile Thr Arg Ile Arg Gly Thr Ser Tyr Gln Ser Pro His Gly Ile Pro Ile Asp Leu Leu Asp

Arg Leu Leu Ile Val Ser Thr Thr Pro Tyr Ser Glu Lys Asp Thr Lys

355 360 365

Gln Ile Leu Arg Ile Arg Cys Glu Glu Glu Asp Val Glu Met Ser Glu 370 375 380

Asp Ala Tyr Thr Val Leu Thr Arg Ile Gly Leu Glu Thr Ser Leu Arg 385 390 395 400

Tyr Ala Ile Gln Leu Ile Thr Ala Ala Ser Leu Val Cys Arg Lys Arg 405 410 415

Lys Gly Thr Glu  $\cdot$ Val Gln Val Asp Asp Ile Lys Arg Val Tyr Ser Leu 420 425 430

Phe Leu Asp Glu Ser Arg Ser Thr Gln Tyr Met Lys Glu Tyr Gln Asp 435 440 445

Ala Phe Leu Phe Asn Glu Leu Lys Gly Glu Thr Met Asp Thr Ser 450 455 460

<210> 7

<211> 3124

<212> PRT

<213> Homo sapiens

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Asn Pro Pro Pro Ser Pro Ala Ala Pro Phe Ala Pro Ser Ala Ser Pro 35 40 45

Ser Ala Pro Gln Ser Pro Ser Tyr Gln Ile Gln Gln Leu Met Asn Arg 50 55 60

Ser Pro Ala Thr Gly Gln Asn Val Asn Ile Thr Leu Gln Ser Val Gly 65 70 75 80

Pro Val Val Gly Gly Asn Gln Gln Ile Thr Leu Ala Pro Leu Pro Leu 85 90 95

Pro Ser Pro Thr Ser Pro Gly Phe Gln Phe Ser Ala Gln Pro Arg Arg
100 105 110

Phe Glu His Gly Ser Pro Ser Tyr Ile Gln Val Thr Ser Pro Leu Ser 115 120 125

Gln Gln Val Gln Thr Gln Ser Pro Thr Gln Pro Ser Pro Gly Pro Gly 130 135 140

Gln Ala Leu Gln Asn Val Arg Ala Gly Ala Pro Gly Pro Gly Leu Gly 145 150 155 160

Leu Cys Ser Ser Ser Pro Thr Gly Gly Phe Val Asp Ala Ser Val Leu 165 170 175

Val Arg Gln Ile Ser Leu Ser Pro Ser Ser Gly Gly His Phe Val Phe 180 185 190

Gln Asp Gly Ser Gly Leu Thr Gln Ile Ala Gln Gly Ala Gln Val Gln 195 200 205

Leu Gln His Pro Gly Thr Pro Ile Thr Val Arg Glu Arg Arg Pro Ser 210 215 220

Gln Pro His Thr Gln Ser Gly Gly Thr Ile His His Leu Gly Pro Gln 225 230 235 240

Ser Pro Ala Ala Gly Gly Ala Gly Leu Gln Pro Leu Ala Ser Pro 245 250 255

Ser His Ile Thr Thr Ala Asn Leu Pro Pro Gln Ile Ser Ser Ile Ile 260 265 270

Gln Gly Gln Leu Val Gln Gln Gln Gln Val Leu Gln Gly Pro Pro Leu 275 280 285

Pro Arg Pro Leu Gly Phe Glu Arg Thr Pro Gly Val Leu Leu Pro Gly 290 295 300

Ala Gly Gly Ala Ala Gly Phe Gly Met Thr Ser Pro Pro Pro Pro Thr 305 310 315 320

Ser Pro Ser Arg Thr Ala Val Pro Pro Gly Leu Ser Ser Leu Pro Leu 325 330 335

Thr Ser Val Gly Asn Thr Gly Met Lys Lys Val Pro Lys Lys Leu Glu 340 345 350

Glu Ile Pro Pro Ala Ser Pro Glu Met Ala Gln Met Arg Lys Gln Cys 355 360 365

Leu Asp Tyr His Tyr Gln Glu Met Gln Ala Leu Lys Glu Val Phe Lys 370 375 380

Glu Tyr Leu Ile Glu Leu Phe Phe Leu Gln His Phe Gln Gly Asn Met 385 390 395 400

Met Asp Phe Leu Ala Phe Lys Lys Lys His Tyr Ala Pro Leu Gln Ala 405 410 415

Tyr Leu Arg Gln Asn Asp Leu Asp Ile Glu Lys Ser Glu Val Ile Asn Asp Glu Gln Gln Ala Leu Ala Gly Ser Leu Val Ala Gly Ala Gly Ser Thr Val Glu Thr Asp Leu Phe Lys Arg Gln Gln Ala Met Pro Ser Thr Gly Met Ala Glu Gln Ser Lys Arg Pro Arg Leu Glu Val Gly His Gln Gly Val Val Phe Gln His Pro Gly Ala Asp Ala Gly Val Pro Leu Gln Gln Leu Met Pro Thr Ala Gln Gly Gly Met Pro Pro Thr Pro Gln Ala Ala Gln Leu Ala Gly Gln Arg Gln Ser Gln Gln Gln Tyr Asp Pro Ser Thr Gly Pro Pro Val 530 535 Gln Asn Ala Ala Ser Leu His Thr Pro Leu Pro Gln Leu Pro Gly Arg 545 Leu Pro Pro Ala Gly Val Pro Thr Ala Ala Leu Ser Ser Ala Leu Gln 570 Phe Ala Gln Gln Pro Gln Val Val Glu Ala Gln Thr Gln Leu Gln Ile Pro Val Lys Thr Gln Gln Pro Asn Val Pro Ile Pro Ala Pro Pro Ser 600 595 Ser Gln Leu Pro Ile Pro Pro Ser Gln Pro Ala Gln Leu Ala Leu His Val Pro Thr Pro Gly Lys Val Gln Val Gln Ala Ser Gln Leu Ser Ser 635 Leu Pro Gln Met Val Ala Ser Thr Arg Leu Pro Val Asp Pro Ala Pro Pro Cys Pro Arg Pro Leu Pro Thr Ser Ser Thr Ser Ser Leu Ala Pro 660 665 Val Ser Gly Ser Gly Pro Gly Pro Ser Pro Ala Arg Ser Ser Pro Val Asn Arg Pro Ser Ser Ala Thr Asn Lys Ala Leu Ser Pro Val Thr Ser 690 695 700

Arg Thr Pro Gly Val Val Ala Ser Ala Pro Thr Lys Pro Gln Ser Pro Ala Gln Asn Ala Thr Ser Ser Gln Asp Ser Ser Gln Asp Thr Leu Thr Glu Gln Ile Thr Leu Glu Asn Gln Val His Gln Arg Ile Ala Glu Leu 745 Arg Lys Ala Gly Leu Trp Ser Gln Arg Arg Leu Pro Lys Leu Gln Glu Ala Pro Arg Pro Lys Ser His Trp Asp Tyr Leu Leu Glu Glu Met Gln Trp Met Ala Thr Asp Phe Ala Gln Glu Arg Arg Trp Lys Val Ala Ala Ala Lys Lys Leu Val Arg Thr Val Val Arg His His Glu Glu Lys Gln 810 805 Leu Arg Glu Glu Arg Gly Lys Lys Glu Glu Gln Ser Arg Leu Arg Arg Ile Ala Ala Ser Thr Ala Arg Glu Ile Glu Cys Phe Trp Ser Asn Ile 840 Glu Gln Val Val Glu Ile Lys Leu Arg Val Glu Leu Glu Glu Lys Arg Lys Lys Ala Leu Asn Leu Gln Lys Val Ser Arg Arg Gly Lys Glu Leu 875 865 870 Arg Pro Lys Gly Phe Asp Ala Leu Gln Glu Ser Ser Leu Asp Ser Gly 890 885 Met Ser Gly Arg Lys Arg Lys Ala Ser Ile Ser Leu Thr Asp Asp Glu 900 905 Val Asp Asp Glu Glu Glu Thr Ile Glu Glu Glu Glu Ala Asn Glu Gly 915 Val Val Asp His Gln Thr Glu Leu Ser Asn Leu Ala Lys Glu Ala Glu 930 935 Leu Pro Leu Leu Asp Leu Met Lys Leu Tyr Glu Gly Ala Phe Leu Pro Ser Ser Gln Trp Pro Arg Pro Lys Pro Asp Gly Glu Asp Thr Ser Gly 970

- Glu Glu Asp Ala Asp Asp Cys Pro Gly Asp Arg Glu Ser Arg Lys Asp 980 985 990
- Leu Val Leu Ile Asp Ser Leu Phe Ile Met Asp Gln Phe Lys Ala Ala 995 1000 1005
- Glu Arg Met Asn Ile Gly Lys Pro Asn Ala Lys Asp Ile Ala Asp 1010 1015 1020
- Val Thr Ala Val Ala Glu Ala Ile Leu Pro Lys Gly Ser Ala Arg 1025 1030 1035
- Val Thr Thr Ser Val Lys Phe Asn Ala Pro Ser Leu Leu Tyr Gly 1040 1045 1050
- Ala Leu Arg Asp Tyr Gln Lys Ile Gly Leu Asp Trp Leu Ala Lys 1055 1060 1065
- Leu Tyr Arg Lys Asn Leu Asn Gly Ile Leu Ala Asp Glu Ala Gly 1070 1075 1080
- Leu Gly Lys Thr Val Gln Ile Ile Ala Phe Phe Ala His Leu Ala 1085 1090 1095
- Cys Asn Glu Gly Asn Trp Gly Pro His Leu Val Val Val Arg Ser 1100 1105 1110
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- Gly Leu Lys Ile Leu Ser Tyr Ile Gly Ser His Arg Glu Leu Lys 1130 1135 1140
- Ala Lys Arg Gln Glu Trp Ala Glu Pro Asn Ser Phe His Val Cys 1145 1150 1155
- Ile Thr Ser Tyr Thr Gln Phe Phe Arg Gly Leu Thr Ala Phe Thr 1160 1165 1170
- Arg Val Arg Trp Lys Cys Leu Val Ile Asp Glu Met Gln Arg Val 1175 1180 1185
- Lys Gly Met Thr Glu Arg His Trp Glu Ala Val Phe Thr Leu Gln 1190 1195 1200
- Ser Gln Gln Arg Leu Leu Ile Asp Ser Pro Leu His Asn Thr 1205 1210 1215
- Phe Leu Glu Leu Trp Thr Met Val His Phe Leu Val Pro Gly Ile 1220 1225 1230

Ser Arg Pro Tyr Leu Ser Ser Pro Leu Arg Ala Pro Ser Glu Glu 1235 1240 1245

Ser Gln Asp Tyr Tyr His Lys Val Val Ile Arg Leu His Arg Val 1250 1255 1260

Thr Gln Pro Phe Ile Leu Arg Arg Thr Lys Arg Asp Val Glu Lys 1265 1270 1275

Gln Leu Thr Lys Lys Tyr Glu His Val Leu Lys Cys Arg Leu Ser 1280 1285 1290

Asn Arg Gln Lys Ala Leu Tyr Glu Asp Val Ile Leu Gln Pro Gly 1295 1300 1305

Thr Gln Glu Ala Leu Lys Ser Gly His Phe Val Asn Val Leu Ser 1310 1315 1320

Ile Leu Val Arg Leu Gln Arg Ile Cys Asn His Pro Gly Leu Val 1325 1330 1335

Glu Pro Arg His Pro Gly Ser Ser Tyr Val Ala Gly Pro Leu Glu 1340 1345 1350

Tyr Pro Ser Ala Ser Leu Ile Leu Lys Ala Leu Glu Arg Asp Phe 1355 1360 1365

Trp Lys Glu Ala Asp Leu Ser Met Phe Asp Leu Ile Gly Leu Glu 1370 1380

Asn Lys Ile Thr Arg His Glu Ala Glu Leu Leu Ser Lys Lys 1385 1390 1395

Ala Ala Arg Pro Ala Ala Ala Lys Leu Lys Ala Ser Arg Leu Phe 1415 1420 1425

Gln Pro Val Gln Tyr Gly Gln Lys Pro Glu Gly Arg Thr Val Ala 1430 1440

Phe Pro Ser Thr His Pro Pro Arg Thr Ala Ala Pro Thr Thr Ala 1445 1450 1455

Ser Ala Ala Pro Gln Gly Pro Leu Arg Gly Arg Pro Pro Ile Ala 1460 1465 1470

Thr Phe Ser Ala Asn Pro Glu Ala Lys Ala Ala Ala Ala Pro Phe 1475 1480 1485

- Gln Thr Ser Gln Ala Ser Ala Ser Ala Pro Arg His Gln Pro Ala 1490 1495 1500
- Ser Ala Ser Ser Thr Ala Ala Ser Pro Ala His Pro Ala Lys Leu 1505 1510 1515
- Arg Ala Gln Thr Thr Ala Gln Ala Phe Thr Pro Gly Gln Pro Pro 1520 1530
- Pro Gln Pro Gln Ala Pro Ser His Ala Ala Gly Gln Ser Ala Leu 1535 1540 1545
- Pro Gln Arg Leu Val Leu Pro Ser Gln Ala Gln Ala Arg Leu Pro 1550 1560
- Ser Gly Glu Val Val Lys Ile Ala Gln Leu Ala Ser Ile Thr Gly 1565 1570 1575
- Pro Gln Ser Arg Val Ala Gln Pro Glu Thr Pro Val Thr Leu Gln 1580 1590
- Phe Gln Gly Ser Lys Phe Thr Leu Ser His Ser Gln Phe Arg Gln 1595 1600 1605
- Phe Thr Ala Gly Gln Pro Leu Gln Leu Gln Gly Ser Val Leu Gln 1610 1620
- Ile Val Ser Ala Pro Gly Gln Pro Tyr Leu Arg Ala Pro Gly Pro 1625 1630 1635
- Val Val Met Gln Thr Val Ser Gln Ala Gly Ala Val His Gly Ala 1640 1650
- Leu Gly Ser Lys Pro Pro Ala Gly Gly Pro Ser Pro Ala Pro Leu 1655 1660 1665
- Thr Pro Gln Val Gly Val Pro Gly Arg Val Ala Val Asn Ala Leu 1670 1680
- Ala Val Gly Glu Pro Gly Thr Ala Ser Lys Pro Ala Ser Pro Ile 1685 1690 1695
- Gly Gly Pro Thr Gln Glu Glu Lys Thr Arg Leu Leu Lys Glu Arg 1700 1705 1710
- Leu Asp Gln Ile Tyr Leu Val Asn Glu Arg Arg Cys Ser Gln Ala 1715 1720 1725
- Pro Val Tyr Gly Arg Asp Leu Leu Arg Ile Cys Ala Leu Pro Ser 1730 1735 1740
- His Gly Arg Val Gln Trp Arg Gly Ser Leu Asp Gly Arg Arg Gly

1745 1750 1755

Lys	Glu 1760		Gly	Pro	Ala	His 1765	Ser	Tyr	Thr	Ser	Ser 1770	Ser	Glu	Ser
Pro	Ser 1775	Glu	Leu	Met	Leu	Thr 1780	Leu	Cys	Arg	Cys	Gly 1785	Glu	Ser	Leu
Gln	Asp 1790	Val	Ile	Asp	Arg	Val 1795	Ala	Phe	Val	Ile	Pro 1800	Pro	Val	Val
Ala	Ala 1805		Pro	Ser	Leu	Arg 1810	Val	Pro	Arg	Pro	Pro 1815	Pro	Leu	Tyr
Ser	His 1820		Met	Arg	Ile	Leu 1825		Gln	Gly	Leu	Arg 1830	Glu	His	Ala
Ala	Pro 1835	Tyr	Phe	Gln	Gln	Leu 1840	Arg	Gln	Thr	Thr	Ala 1845	Pro	Arg	Leu
Leu	Gln 1850		Pro	Glu	Leu	Arg 1855	Leu	Val	Gln	Phe	Asp 1860	Ser	Gly	Lys
Leu	Glu 1865	Ala	Leu	Ala	Ile	Leu 1870	Leu	Gln	Lys	Leu	Lys 1875		Glu	Gly
Arg	Arg 1880		Leu	Ile	Leu	Ser 1885		Met	Ile	Leu	Met 1890	Leu	Asp	Ile
Leu	Glu 1895	Met	Phe	Leu	Asn	Phe 1900	His	Tyr	Leu	Thr	Tyr 1905	Val	Arg	Ile
Asp	Glu 1910	Asn	Ala	Ser	Ser	Glu 1915		Arg	Gln	Glu	Leu 1920	Met	Arg	Ser
Phe	Asn 1925	Arg	Asp	Arg	Arg	Ile 1930	Phe	Cys	Ala	Ile	Leu 1935	Ser	Thr	His
Ser	Arg 1940	Thr	Thr	Gly	Ile	Asn 1945	Leu	Val	Glu	Ala	Asp 1950	Thr	Val	Val
Phe	Tyr 1955		Asn	Asp	Leu	Asn 1960		Val	Met	Asp	Ala 1965	Lys	Ala	Gln
Glu	Trp 1970		Asp	Arg	Ile	Gly 1975	Arg	Cys	Lys	Asp	Ile 1980	His	Ile	Tyr
Arg	Leu 1985		Ser	Gly	Asn	Ser 1990		Glu	Glu	Lys	Leu 1995	Leu	Lys	Asn
Gly	Thr 2000		Asp	Leu	Ile	Arg 2005	Glu	Val	Ala	Ala	Gln 2010	Gly	Asn	Asp

Tyr Ser Met Ala Phe Leu Thr Gln Arg Thr Ile Gln Glu Leu Phe 2015 2020 2025

Glu Val Tyr Ser Pro Met Asp Asp Ala Gly Phe Pro Val Lys Ala 2030 2035 2040

Glu Glu Phe Val Val Leu Ser Gln Glu Pro Ser Val Thr Glu Thr 2045 2050 2055

Ile Ala Pro Lys Ile Ala Arg Pro Phe Ile Glu Ala Leu Lys Ser 2060 2065 2070

Ile Glu Tyr Leu Glu Glu Asp Ala Gln Lys Ser Ala Gln Glu Gly 2075 2080 2085

Val Leu Gly Pro His Thr Asp Ala Leu Ser Ser Asp Ser Glu Asn 2090 2095 2100

Met Pro Cys Asp Glu Glu Pro Ser Gln Leu Glu Glu Leu Ala Asp 2105 2110 2115

Phe Met Glu Gln Leu Thr Pro Ile Glu Lys Tyr Ala Leu Asn Tyr 2120 2125 2130

Leu Glu Leu Phe His Thr Ser Ile Glu Glu Glu Lys Glu Arg Asn 2135 2140 2145

Ser Glu Asp Ala Val Met Thr Ala Val Arg Ala Trp Glu Phe Trp 2150 2160

Asn Leu Lys Thr Leu Gln Glu Arg Glu Ala Arg Leu Arg Leu Glu 2165 2170 2175

Gln Glu Glu Ala Glu Leu Leu Thr Tyr Thr Arg Glu Asp Ala Tyr 2180 2185 2190

Ser Met Glu Tyr Val Tyr Glu Asp Val Asp Gly Gln Thr Glu Val 2195 . 2200 2205

Met Pro Leu Trp Thr Pro Pro Thr Pro Pro Gln Asp Asp Ser Asp 2210 2215 2220

Ile Tyr Leu Asp Ser Val Met Cys Leu Met Tyr Glu Ala Thr Pro 2225 2230 2235

Ile Pro Glu Ala Lys Leu Pro Pro Val Tyr Val Arg Lys Glu Arg 2240 2245 2250

Lys Arg His Lys Thr Asp Pro Ser Ala Ala Gly Arg Lys Lys 2255 2260 2265

Gln Arg His Gly Glu Ala Val Val Pro Pro Arg Ser Leu Phe Asp 2270 2275 2280

Arg Ala Thr Pro Gly Leu Leu Lys Ile Arg Arg Glu Gly Lys Glu 2285 2290 2295

Gln Lys Lys Asn Ile Leu Leu Lys Gln Gln Val Pro Phe Ala Lys 2300 2305 2310

Pro Leu Pro Thr Phe Ala Lys Pro Thr Ala Glu Pro Gly Gln Asp 2315 2320 2325

Asn Pro Glu Trp Leu Ile Ser Glu Asp Trp Ala Leu Leu Gln Ala 2330 2340

Val Lys Gln Leu Leu Glu Leu Pro Leu Asn Leu Thr Ile Val Ser 2345 2350 2355

Pro Ala His Thr Pro Asn Trp Asp Leu Val Ser Asp Val Val Asn 2360 2365 2370

Ser Cys Ser Arg Ile Tyr Arg Ser Ser Lys Gln Cys Arg Asn Arg 2375 2380 2385

Tyr Glu Asn Val Ile Ile Pro Arg Glu Glu Gly Lys Ser Lys Asn 2390 2395 2400

Asn Arg Pro Leu Arg Thr Ser Gln Ile Tyr Ala Gln Asp Glu Asn 2405 2410 2415

Ala Thr His Thr Gln Leu Tyr Thr Ser His Phe Asp Leu Met Lys 2420 2425 2430

Met Thr Ala Gly Lys Arg Ser Pro Pro Ile Lys Pro Leu Leu Gly 2435 2440 2445

Met Asn Pro Phe Gln Lys Asn Pro Lys His Ala Ser Val Leu Ala 2450 2455 2460

Glu Ser Gly Ile Asn Tyr Asp Lys Pro Leu Pro Pro Ile Gln Val 2465 2470 2475

Ala Ser Leu Arg Ala Glu Arg Ile Ala Lys Glu Lys Lys Ala Leu 2480 2485 2490

Ala Asp Gln Gln Lys Ala Gln Gln Pro Ala Val Ala Gln Pro Pro 2495 2500 2505

Pro Pro Gln Pro Gln Pro Pro Pro Pro Gln Gln Pro Pro Pro 2510 2515 2520

Pro Leu Pro Gln Pro Gln Ala Ala Gly Ser Gln Pro Pro Ala Gly 2525 2530 2535

Pro Pro Ala Val Gln Pro Gln Pro Gln Pro Gln Pro Gln Thr Gln 2540 2545 2550

Pro Gln Pro Val Gln Ala Pro Ala Lys Ala Gln Pro Ala Ile Thr 2555 2560 2565

Thr Gly Gly Ser Ala Ala Val Leu Ala Gly Thr Ile Lys Thr Ser 2570 2580

Val Thr Gly Thr Ser Met Pro Thr Gly Ala Val Ser Gly Asn Val 2585 2590 2595

Ile Val Asn Thr Ile Ala Gly Val Pro Ala Ala Thr Phe Gln Ser 2600 2605 2610

Ile Asn Lys Arg Leu Ala Ser Pro Val Ala Pro Gly Ala Leu Thr 2615 2620 2625

Thr Pro Gly Gly Ser Ala Pro Ala Gln Val Val His Thr Gln Pro 2630  $\phantom{0}2635$   $\phantom{0}2640$ 

Pro Pro Arg Ala Val Gly Ser Pro Ala Thr Ala Thr Pro Asp Leu 2645 2655

Val Ser Met Ala Thr Thr Gln Gly Val Arg Ala Val Thr Ser Val 2660 2670

Thr Ala Ser Ala Val Val Thr Thr Asn Leu Thr Pro Val Gln Thr 2675 2680 2685

Pro Ala Arg Ser Leu Val Pro Gln Val Ser Gln Ala Thr Gly Val 2690 2695 2700

Gln Leu Pro Gly Lys Thr Ile Thr Pro Ala His Phe Gln Leu Leu 2705 2710 2715

Gln Thr Thr Thr Ser Gln Val Gln Val Pro Gln Ile Gln Gly 2750 2755 2760

Gln Ala Gln Ser Pro Ala Gln Ile Lys Ala Val Gly Lys Leu Thr 2765 2770 2775

Pro Glu His Leu Ile Lys Met Gln Lys Gln Lys Leu Gln Met Pro

2780 2785 2790

Pro	Gln 2795	Pro	Pro	Pro	Pro	Gln 2800	Ala	Gln	Ser	Ala	Pro 2805	Pro	Gln	Pro
Ala	Ala 2810	Gln	Val	Gln		Gln 2815	Thr	Ser	Gln	Pro	Pro 2820	Gln	Gln	Gln
Ser	Pro 2825	Gln	Leu	Thr	Thr	Val 2830	Thr	Ala	Pro	Arg	Pro 2835	Gly	Ala	Leu
Leu	Thr 2840	Gly	Thr	Thr	Val	Ala 2845	Asn	Leu	Gln	Val	Ala 2850	Arg	Leu	Thr
Arg	Val 2855	Pro	Thr	Ser		Leu 2860		Ser	Gln	Gly	Gln 2865		Gln	Thr
Gln	Ala 2870	Pro	Gln	Pro	Ala	Gln 2875	Val	Pro	Leu	Pro	Lys 2880	Pro	Pro	Val
Val	Ser 2885	Val	Pro	Ala	Ala	Val 2890	Val	Ser	Ser	Pro	Gly 2895	Val	Thr	Thr
Leu	Pro 2900	Met	Asn	Val	Ala	Gly 2905	Ile	Ser	Val	Ala	Ile 2910		Gln	Pro
Gln	Lys 2915		Ala	Gly	Gln	Thr 2920	Val	Val	Ala	Gln	Pro 2925	Val	His	Met
Gln	Gln 2930	Leu	Leu	Lys	Leu	Lys 2935		Gln	Ala	Val	Gln 2940		Gln	Lys
Ala	Ile 2945	Gln	Pro	Gln	Ala	Ala 2950	Gln	Gly	Pro	Ala	Thr 2955	Val	Gln	Gln
Lys	Ile 2960	Thr	Ala	Gln	Gln	Ile 2965	Thr	Thr	Pro	Gly	Ala 2970	Gln	Gln	Lys
Val	Ala 2975	Tyr	Ala	Ala	Gln	Pro 2980	Ala	Leu	Lys	Thr	Gln 2985	Phe	Leu	Thr
Thr	Pro 2990	Ile	Ser	Gln	Ala	Gln 2995	Lys	Leu	Ala	Gly	Ala 3000	Gln	Gln	Val
Gln	Thr 3005	Gln	Ile	Gln	Val	Ala 3010	Lys	Leu	Pro	Gln	Val 3015		Gln	Gln
Gln	Thr 3020	Pro	Val	Ala	Ser	Ile 3025	Gln	Gln	Val	Ala	Ser 3030	Ala	Ser	Gln
Gln	Ala 3035	Ser	Pro	Gln	Thr	Val 3040	Ala	Leu	Thr	Gln	Ala 3045	Thr	Ala	Ala

Gly Gln Gln Val Gln Met Ile Pro Ala Val Thr Ala Thr Ala Gln 3050 3055 3060

Val Val Gln Gln Lys Leu Ile Gln Gln Gln Val Val Thr Thr Ala 3065 3070 3075

Ser Ala Pro Leu Gln Thr Pro Gly Ala Pro Asn Pro Ala Gln Val 3080 3085 3090

Pro Ala Ser Ser Asp Ser Pro Ser Gln Gln Pro Lys Leu Gln Met 3095 3100 3105

Arg Val Pro Ala Val Arg Leu Lys Thr Pro Thr Lys Pro Pro Cys 3110 3115 3120

Gln

<210> 8

<211> 836

<212> PRT

<213> Homo sapiens

<400> 8

Met Ser Lys Leu Ser Phe Arg Ala Arg Ala Leu Asp Ala Ser Lys Pro 1 10 15

Leu Pro Val Phe Arg Cys Glu Asp Leu Pro Asp Leu His Glu Tyr Ala 20 25 30

Ser Ile Asn Arg Ala Val Pro Gln Met Pro Thr Gly Met Glu Lys Glu 35 40 45

Glu Glu Ser Glu His His Leu Gln Arg Ala Ile Ser Ala Gln Gln Val 50 55 60

Tyr Gly Glu Lys Arg Asp Asn Met Val Ile Pro Val Pro Glu Ala Glu 65 70 75 80

Ser Asn Ile Ala Tyr Tyr Glu Ser Ile Tyr Pro Gly Glu Phe Lys Met 85 90 95

Pro Lys Gln Leu Ile His Ile Gln Pro Phe Ser Leu Asp Ala Glu Gln 100 105 110

Pro Asp Tyr Asp Leu Asp Ser Glu Asp Glu Val Phe Val Asn Lys Leu

Lys Lys Lys Met Asp Ile Cys Pro Leu Gln Phe Glu Glu Met Ile Asp 130 135 140

Arg Leu Glu Lys Gly Ser Gly Gln Gln Pro Val Ser Leu Gln Glu Ala Lys Leu Leu Lys Glu Asp Asp Glu Leu Ile Arg Glu Val Tyr Glu Tyr Trp Ile Lys Lys Arg Lys Asn Cys Arg Gly Pro Ser Leu Ile Pro Ser Val Lys Gln Glu Lys Arg Asp Gly Ser Ser Thr Asn Asp Pro Tyr Val Ala Phe Arg Arg Arg Thr Glu Lys Met Gln Thr Arg Lys Asn Arg Lys Asn Asp Glu Ala Ser Tyr Glu Lys Met Leu Lys Leu Arg Arg Asp Leu Ser Arg Ala Val Thr Ile Leu Glu Met Ile Lys Arg Arg Glu Lys Ser Lys Arg Glu Leu Leu His Leu Thr Leu Glu Ile Met Glu Lys Arg 260 Tyr Asn Leu Gly Asp Tyr Asn Gly Glu Ile Met Ser Glu Val Met Ala Gln Arg Gln Pro Met Lys Pro Thr Tyr Ala Ile Pro Ile Ile Pro Ile Thr Asn Ser Ser Gln Phe Lys His Gln Glu Ala Met Asp Val Lys Glu 305 Phe Lys Val Asn Lys Gln Asp Lys Ala Asp Leu Ile Arg Pro Lys Arg Lys Tyr Glu Lys Lys Pro Lys Val Leu Pro Ser Ser Ala Ala Ala Thr Pro Gln Gln Thr Ser Pro Ala Ala Leu Pro Val Phe Asn Ala Lys Asp Leu Asn Gln Tyr Asp Phe Pro Ser Ser Asp Glu Glu Pro Leu Ser Gln Val Leu Ser Gly Ser Ser Glu Ala Glu Glu Asp Asn Asp Pro Asp Gly 400 Pro Phe Ala Phe Arg Arg Lys Ala Gly Cys Gln Tyr Tyr Ala Pro His 405 415 Leu Asp Gln Thr Gly Asn Trp Pro Trp Thr Ser Pro Lys Asp Gly Gly Leu Gly Asp Val Arg Tyr Arg Tyr Cys Leu Thr Thr Leu Thr Val Pro 435 440 Gln Arg Cys Ile Gly Phe Ala Arg Arg Arg Val Gly Arg Gly Gly Arg Val Leu Leu Asp Arg Ala His Ser Asp Tyr Asp Ser Val Phe His His Leu Asp Leu Glu Met Leu Ser Ser Pro Gln His Ser Pro Val Asn Gln 490 Phe Ala Asn Thr Ser Glu Thr Asn Thr Ser Asp Lys Ser Phe Ser Lys 505 Asp Leu Ser Gln Ile Leu Val Asn Ile Lys Ser Cys Arg Trp Arg His Phe Arg Pro Arg Thr Pro Ser Leu His Asp Ser Asp Asn Asp Glu Leu 535 Ser Cys Arg Lys Leu Tyr Arg Ser Ile Asn Arg Thr Gly Thr Ala Gln Pro Gly Thr Gln Thr Cys Ser Thr Ser Thr Gln Ser Lys Ser Ser Ser 570 Gly Ser Ala His Phe Ala Phe Thr Ala Glu Gln Tyr Gln Gln His Gln Gln Gln Leu Ala Leu Met Gln Lys Gln Gln Leu Ala Gln Ile Gln Gln 595 600 Gln Gln Ala Asn Ser Asn Ser Ser Thr Asn Thr Ser Gln Asn Leu Ala 615 610 Ser Asn Gln Gln Lys Ser Gly Phe Arg Leu Asn Ile Gln Gly Leu Glu 625 Arg Thr Leu Gln Gly Phe Val Ser Lys Thr Leu Asp Ser Ala Ser Ala Gln Phe Ala Ala Ser Ala Leu Val Thr Ser Glu Gln Leu Met Gly Phe Lys Met Lys Asp Asp Val Val Leu Gly Ile Gly Val Asn Gly Val Leu Pro Ala Ser Gly Val Tyr Lys Gly Leu His Leu Ser Ser Thr Thr Pro

695

Thr Ala Leu Val His Thr Ser Pro Ser Thr Ala Gly Ser Ala Leu Leu 705 710 715 720

Gln Pro Ser Asn Ile Thr Gln Thr Ser Ser Ser His Ser Ala Leu Ser 725 730 735

His Gln Val Thr Ala Ala Asn Ser Ala Thr Thr Gln Val Leu Ile Gly 740 745 750

Asn Asn Ile Arg Leu Thr Val Pro Ser Ser Val Ala Thr Val Asn Ser 755 760 765

Ile Ala Pro Ile Asn Ala Arg His Ile Pro Arg Thr Leu Ser Ala Val 770 775 780

Pro Ser Ser Ala Leu Lys Leu Ala Ala Ala Ala Asn Cys Gln Val Ser 785 790 795 800

Lys Val Pro Ser Ser Ser Ser Val Asp Ser Val Pro Arg Glu Asn His 805 810 815

Glu Ser Glu Lys Pro Ala Leu Asn Asn Ile Ala Asp Asn Thr Val Ala 820 825 830

Met Glu Val Thr 835

<210> 9

<211> 227

<212> PRT

<213> Homo sapiens

<400> 9

Met Phe Lys Arg Met Ala Glu Phe Gly Pro Asp Ser Gly Gly Arg Val  $1 \hspace{1cm} 5 \hspace{1cm} 10 \hspace{1cm} 15$ 

Lys Gly Val Thr Ile Val Lys Pro Ile Val Tyr Gly Asn Val Ala Arg 20 25 30

Tyr Phe Gly Lys Lys Arg Glu Glu Asp Gly His Thr His Gln Trp Thr 35 40 45

Val Tyr Val Lys Pro Tyr Arg Asn Glu Asp Met Ser Ala Tyr Val Lys 50 60

Lys Ile Gln Phe Lys Leu His Glu Ser Tyr Gly Asn Pro Leu Arg Val 65 70 75 80

Val Thr Lys Pro Pro Tyr Glu Ile Thr Glu Thr Gly Trp Gly Glu Phe 85 90 95

Glu Ile Ile Ile Lys Ile Phe Phe Ile Asp Pro Asn Glu Arg Pro Val

Thr Leu Tyr His Leu Leu Lys Leu Phe Gln Ser Asp Thr Asn Ala Met 115 120 125

Leu Gly Lys Lys Thr Val Val Ser Glu Phe Tyr Asp Glu Met Ile Phe 130 135 140

Gln Asp Pro Thr Ala Met Met Gln Gln Leu Leu Thr Thr Ser Arg Gln 145 150 155 160

Leu Thr Leu Gly Ala Tyr Lys His Glu Thr Glu Phe Ala Glu Leu Glu 165 170 175

Val Lys Thr Arg Glu Lys Leu Glu Ala Ala Lys Lys Lys Thr Ser Phe 180 185 190

Glu Ile Ala Glu Leu Lys Glu Arg Leu Lys Ala Ser Arg Glu Thr Ile 195 200 205

Asn Cys Leu Lys Asn Glu Ile Arg Lys Leu Glu Glu Asp Asp Gln Ala 210 215 220

Lys Asp Ile 225

<210> 10

<211> 482

<212> PRT

<213> Homo sapiens

<400> 10

Met Ala Gln Thr Gln Gly Thr Arg Arg Lys Val Cys Tyr Tyr Tyr Asp 1 10 15

Gly Asp Val Gly Asn Tyr Tyr Tyr Gly Gln Gly His Pro Met Lys Pro 20 25 30

His Arg Ile Arg Met Thr His Asn Leu Leu Leu Asn Tyr Gly Leu Tyr 35 40 45

Arg Lys Met Glu Ile Tyr Arg Pro His Lys Ala Asn Ala Glu Glu Met 50 60

Thr Lys Tyr His Ser Asp Asp Tyr Ile Lys Phe Leu Arg Ser Ile Arg 65 70 75 80

Pro Asp Asn Met Ser Glu Tyr Ser Lys Gln Met Gln Arg Phe Asn Val

Gly Glu Asp Cys Pro Val Phe Asp Gly Leu Phe Glu Phe Cys Gln Leu 100 105 110 Ser Thr Gly Gly Ser Val Ala Ser Ala Val Lys Leu Asn Lys Gln Gln 115 120 125

Thr Asp Ile Ala Val Asn Trp Ala Gly Gly Leu His His Ala Lys Lys 130 135 140

Ser Glu Ala Ser Gly Phe Cys Tyr Val Asn Asp Ile Val Leu Ala Ile 145 150 155 160

Leu Glu Leu Leu Lys Tyr His Gln Arg Val Leu Tyr Ile Asp Ile Asp 165 170 175

Ile His His Gly Asp Gly Val Glu Glu Ala Phe Tyr Thr Asp Arg
180 185 190

Val Met Thr Val Ser Phe His Lys Tyr Gly Glu Tyr Phe Pro Gly Thr 195 200 205

Gly Asp Leu Arg Asp Ile Gly Ala Gly Lys Gly Lys Tyr Tyr Ala Val 210 215 220

Asn Tyr Pro Leu Arg Asp Gly Ile Asp Asp Glu Ser Tyr Glu Ala Ile 225 230 235 240

Phe Lys Pro Val Met Ser Lys Val Met Glu Met Phe Gln Pro Ser Ala 245 250 255

Val Val Leu Gln Cys Gly Ser Asp Ser Leu Ser Gly Asp Arg Leu Gly 260 265 270

Cys Phe Asn Leu Thr Ile Lys Gly His Ala Lys Cys Val Glu Phe Val 275 280 285

Lys Ser Phe Asn Leu Pro Met Leu Met Leu Gly Gly Gly Tyr Thr 290 295 300

Ile Arg Asn Val Ala Arg Cys Trp Thr Tyr Glu Thr Ala Val Ala Leu 305 310 315 320

Asp Thr Glu Ile Pro Asn Glu Leu Pro Tyr Asn Asp Tyr Phe Glu Tyr 325 330 335

Phe Gly Pro Asp Phe Lys Leu His Ile Ser Pro Ser Asn Met Thr Asn 340 345 350

Gln Asn Thr Asn Glu Tyr Leu Glu Lys Ile Lys Gln Arg Leu Phe Glu 355 365

Asn Leu Arg Met Leu Pro His Ala Pro Gly Val Gln Met Gln Ala Ile 370 375 380

Pro Glu Asp Ala Ile Pro Glu Glu Ser Gly Asp Glu Asp Glu Asp Asp 385 390 395 400

Pro Asp Lys Arg Ile Ser Ile Cys Ser Ser Asp Lys Arg Ile Ala Cys 405 410 415

Glu Glu Glu Phe Ser Asp Ser Glu Glu Glu Gly Gly Gly Arg Lys 420 425 430

Asn Ser Ser Asn Phe Lys Lys Ala Lys Arg Val Lys Thr Glu Asp Glu 435 440 445

Lys Glu Lys Asp Pro Glu Glu Lys Lys Glu Val Thr Glu Glu Glu Lys 450 455 460

Thr Lys Glu Glu Lys Pro Glu Ala Lys Gly Val Lys Glu Glu Val Lys 465 470 475 480

Leu Ala

<210> 11

<211> 1727

<212> PRT

<213> Homo sapiens

<400> 11

Ala Leu Leu His Phe Met Arg Glu Lys Glu Gln Glu Arg Glu Gln 1 5 10 15

Leu Met Glu Asp Lys Lys Arg Lys Lys Glu Asp Lys Lys Lys Glu 20 25 30

Ala Thr Gln Lys Val Thr Glu Gln Lys Thr Lys Val Pro Glu Val Thr 35 40 45

Lys Pro Ser Leu Ser Gln Pro Thr Ala Ala Ser Pro Ile Gly Ser Ser 50 55 60

Pro Ser Pro Pro Val Asn Gly Gly Asn Asn Ala Lys Arg Val Ala Val 65 70 75 80

Pro Asn Gly Gln Pro Pro Ser Ala Ala Arg Tyr Met Pro Arg Glu Val 85 90 95

Pro Pro Arg Phe Arg Cys Gln Gln Asp His Lys Val Leu Leu Lys Arg 100 105 110

Gly Gln Pro Pro Pro Pro Ser Cys Met Leu Leu Gly Gly Gly Ala Gly 115 120 125

Pro Pro Pro Cys Thr Ala Pro Gly Ala Asn Pro Asn Asn Ala Gln Val

130 135 140

Thr Gly Ala Leu Leu Gln Ser Glu Ser Gly Thr Ala Pro Asp Ser Thr 150 Leu Gly Gly Ala Ala Ala Ser Asn Tyr Ala Asn Ser Thr Trp Gly Ser Gly Ala Ser Ser Asn Asn Gly Thr Ser Pro Asn Pro Ile His Ile Trp 185 180 Asp Lys Val Ile Val Asp Gly Ser Asp Met Glu Glu Trp Pro Cys Ile Ala Ser Lys Asp Thr Glu Ser Ser Ser Glu Asn Thr Thr Asp Asn Asn 210 Ser Ala Ser Asn Pro Gly Ser Glu Lys Ser Thr Leu Pro Gly Ser Thr Thr Ser Asn Lys Gly Lys Gly Ser Gln Cys Gln Ser Ala Ser Ser Gly Asn Glu Cys Asn Leu Gly Val Trp Lys Ser Asp Pro Lys Ala Lys Ser Val Gln Ser Ser Asn Ser Thr Thr Glu Asn Asn Gly Leu Gly Asn Trp Arg Asn Val Ser Gly Gln Asp Arg Ile Gly Pro Gly Ser Gly Phe Ser Asn Phe Asn Pro Asn Ser Asn Pro Ser Ala Trp Pro Ala Leu Val 315 305 310 Gln Glu Gly Thr Ser Arg Lys Gly Ala Leu Glu Thr Asp Asn Ser Asn Ser Ser Ala Gln Val Ser Thr Val Gly Gln Thr Ser Arg Glu Gln Gln 340 Ser Lys Met Glu Asn Ala Gly Val Asn Phe Val Val Ser Gly Arg Glu Gln Ala Gln Ile His Asn Thr Asp Gly Pro Lys Asn Gly Asn Thr Asn Ser Leu Asn Leu Ser Ser Pro Asn Pro Met Glu Asn Lys Gly Met Pro Phe Gly Met Gly Leu Gly Asn Thr Ser Arg Ser Thr Asp Ala Pro Ser Gln Ser Thr Gly Asp Arg Lys Thr Gly Ser Val Gly Ser Trp Gly Ala 420 425 430

Ala Arg Gly Pro Ser Gly Thr Asp Thr Val Ser Gly Gln Ser Asn Ser 435 440 445

Gly Asn Asn Gly Asn Asn Gly Lys Glu Arg Glu Asp Ser Trp Lys Gly 450 455 460

Ala Ser Val Gln Lys Ser Thr Gly Ser Lys Asn Asp Ser Trp Asp Asn 465 470 475 480

Asn Asn Arg Ser Thr Gly Gly Ser Trp Asn Phe Gly Pro Gln Asp Ser 485 490 495

Asn Asp Asn Lys Trp Gly Glu Gly Asn Lys Met Thr Ser Gly Val Ser 500 505 510

Gln Gly Glu Trp Lys Gln Pro Thr Gly Ser Asp Glu Leu Lys Ile Gly 515 520 525

Glu Trp Ser Gly Pro Asn Gln Pro Asn Ser Ser Thr Gly Ala Trp Asp 530 540

Asn Gln Lys Gly His Pro Leu Pro Glu Asn Gln Gly Asn Ala Gln Ala 545 550 560

Pro Cys Trp Gly Arg Ser Ser Ser Ser Thr Gly Ser Glu Val Gly Gly 565 570 575

Gln Ser Thr Gly Ser Asn His Lys Ala Gly Ser Ser Asp Ser His Asn 580 585 590

Ser Gly Arg Arg Ser Tyr Arg Pro Thr His Pro Asp Cys Gln Ala Val 595 600 605

Leu Gln Thr Leu Leu Ser Arg Thr Asp Leu Asp Pro Arg Val Leu Ser 610 620

Asn Thr Gly Trp Gly Gln Thr Gln Ile Lys Gln Asp Thr Val Trp Asp 625 630 635 640

Ile Glu Glu Val Pro Arg Pro Glu Gly Lys Ser Asp Lys Gly Thr Glu 645  $\phantom{000}650$   $\phantom{000}655$ 

Gly Trp Glu Ser Ala Ala Thr Gln Thr Lys Asn Ser Gly Gly Trp Gly 660 665 670

Asp Ala Pro Ser Gln Ser Asn Gln Met Lys Ser Gly Trp Gly Glu Leu 675 680 685

Ser Ala Ser Thr Glu Trp Lys Asp Pro Lys Asn Thr Gly Gly Trp Asn 690 695 700

Asp Tyr Lys Asn Asn Asn Ser Ser Asn Trp Gly Gly Gly Arg Pro Asp 705 710 715 720

Glu Lys Thr Pro Ser Ser Trp Asn Glu Asn Pro Ser Lys Asp Gln Gly 725 730 735

Trp Gly Gly Gly Arg Gln Pro Asn Gln Gly Trp Ser Ser Gly Lys Asn 740 745 750

Gly Trp Gly Glu Glu Val Asp Gln Thr Lys Asn Ser Asn Trp Glu Ser 755 760 765

Ser Ala Ser Lys Pro Val Ser Gly Trp Gly Glu Gly Gln Asn Glu 770 775 780

Ile Gly Thr Trp Gly Asn Gly Gly Asn Ala Ser Leu Ala Ser Lys Gly 785 . 790 795 800

Gly Trp Glu Asp Cys Lys Arg Ser Pro Ala Trp Asn Glu Thr Gly Arg 805 810 815

Gln Pro Asn Ser Trp Asn Lys Gln His Gln Gln Gln Pro Pro Gln 820 825 830

Gln Pro Pro Pro Gln Pro Glu Ala Ser Gly Ser Trp Gly Gly Pro 835 840 845

Pro Pro Pro Pro Gly Asn Val Arg Pro Ser Asn Ser Ser Trp Ser 850 855 860

Ser Gly Pro Gln Pro Ala Thr Pro Lys Asp Glu Glu Pro Ser Gly Trp 865 870 875 880

Glu Glu Pro Ser Pro Gln Ser Ile Ser Arg Lys Met Asp Ile Asp Asp 885 890 895

Gly Thr Ser Ala Trp Gly Asp Pro Asn Ser Tyr Asn Tyr Lys Asn Val $900 \hspace{1.5cm} 905 \hspace{1.5cm} 910$ 

Asn Leu Trp Asp Lys Asn Ser Gln Gly Gly Pro Ala Pro Arg Glu Pro 915 920 925

Asn Leu Pro Thr Pro Met Thr Ser Lys Ser Ala Ser Asp Ser Lys Ser 930 935 940

Met Gln Asp Gly Trp Gly Glu Ser Asp Gly Pro Val Thr Gly Ala Arg 945 950 955 960

- His Pro Ser Trp Glu Glu Glu Glu Asp Gly Gly Val Trp Asn Thr Thr 965 970 975
- Gly Ser Gln Gly Ser Ala Ser Ser His Asn Ser Ala Ser Trp Gly Gln 980 985 990
- Gly Gly Lys Cys Gln Met Lys Cys Ser Leu Lys Gly Gly Asn Asn Asp 995 1000 1005
- Ser Trp Met Asn Pro Leu Ala Lys Gln Phe Ser Asn Met Gly Leu 1010 1015 1020
- Leu Ser Gln Thr Glu Asp Asn Pro Ser Ser Lys Met Asp Leu Ser 1025 1030 1035
- Val Gly Ser Leu Ser Asp Lys Lys Phe Asp Val Asp Lys Arg Ala 1040 1045 1050
- Met Asn Leu Gly Asp Phe Asn Asp Ile Met Arg Lys Asp Arg Ser 1055 1060 1065
- Gly Phe Arg Pro Pro Asn Ser Lys Asp Met Gly Thr Thr Asp Ser 1070 1075 1080
- Gly Pro Tyr Phe Glu Lys Gly Gly Ser His Gly Leu Phe Gly Asn 1085 1090 1095
- Ser Thr Ala Gln Ser Arg Gly Leu His Thr Pro Val Gln Pro Leu 1100 1105 1110
- Asn Ser Ser Pro Ser Leu Arg Ala Gln Val Pro Pro Gln Phe Ile 1115 1120 1125
- Ser Pro Gln Val Ser Ala Ser Met Leu Lys Gln Phe Pro Asn Ser 1130 1135 1140
- Gly Leu Ser Pro Gly Leu Phe Asn Val Gly Pro Gln Leu Ser Pro 1145 1150 1155
- Gln Gln Ile Ala Met Leu Ser Gln Leu Pro Gln Ile Pro Gln Phe 1160 1165 1170
- Gln Leu Ala Cys Gln Leu Leu Gln Gln Gln Gln Gln Gln Gln 1175 1180 1185
- Leu Leu Gln Asn Gln Arg Lys Ile Ser Gln Ala Val Arg Gln Gln 1190 1195 1200
- Gln Glu Gln Gln Leu Ala Arg Met Val Ser Ala Leu Gln Gln Gln 1205 1210 1215
- Gln Gln Gln Arg Gln Pro Gly Met Lys His Ser Pro Ser His

1220 1225 1230

Pro	Val 1235	Gly	Pro	Lys	Pro	His 1240	Leu	Asp	Asn	Met	Val 1245	Pro	Asn	Ala
Leu	Asn 1250	Val	Gly	Leu	Pro	Asp 1255	Leu	Gln	Thr	Lys	Gly 1260	Pro	Ile	Pro
Gly	Tyr 1265	_	Ser	Gly	Phe	Ser 1270	Ser	Gly	Gly	Met	Asp 1275	Tyr	Gly	Met
Val	Gly 1280	Gly	Lys	Glu	Ala	Gly 1285	Thr	Glu	Ser	Arg	Phe 1290	Lys	Gln	Trp
Thr	Ser 1295	Met	Met	Glu	Gly	Leu 1300	Pro	Ser	Val	Ala	Thr 1305	Gln	Glu	Ala
Asn	Met 1310	His	Lys	Asn	Gly	Ala 1315	Ile	Val	Ala	Pro	Gly 1320	Lys	Thr	Arg
Gly	Gly 1325		Pro	Tyr	Asn	Gln 1330	Phe	Asp	Ile	Ile	Pro 1335	Gly	Asp	Thr
Leu	Gly 1340	Gly	His	Thr	Gly	Pro 1345	Ala	Gly	Asp	Ser	Trp 1350	Leu	Pro	Ala
Lys	Ser 1355	Pro	Pro	Thr	Asn	Lys 1360	Ile	Gly	Ser	Lys	Ser 1365	Ser	Asn	Ala
Ser	Trp 1370		Pro	Glu	Phe	Gln 1375	Pro	Gly	Val	Pro	Trp 1380	Lys	Gly	Ile
Gln	Asn 1385	Ile	Asp	Pro	Glu	Ser 1390	Asp	Pro	Tyr	Val	Thr 1395	Pro	Gly	Ser
Val	Leu 1400	Gly	Gly	Thr	Ala	Thr 1405	Ser	Pro	Ile	Val	Asp 1410	Thr	Asp	His
Gln	Leu 1415	Leu	Arg	Asp	Asn	Thr 1420	Thr	Gly	Ser	Asn	Ser 1425	Ser	Leu	Asn
Thr	Ser 1430	Leu	Pro	Ser	Pro	Gly 1435	Ala	Trp	Pro	Tyr	Ser 1440	Ala	Ser	Asp
Asn	Ser 1445	Phe	Thr	Asn	Val	His 1 <sup>.</sup> 450		Thr	Ser	Ala	Lys 1455		Pro	Asp
Tyr	Lys 1460		Thr	Trp	Ser	Pro 1465	Asp	Pro	Ile	Gly	His 1470	Asn	Pro	Thr
His	Leu 1475		Asn	Lys	Met	Trp 1480		Asn	His	Ile	Ser 1485		Arg	Asn

Thr Thr Pro Leu Pro Arg Pro Pro Pro Gly Leu Thr Asn Pro Lys 1495 1490 Pro Ser Ser Pro Trp Ser Ser Thr Ala Pro Arg Ser Val Arg Gly 1510 Trp Gly Thr Gln Asp Ser Arg Leu Ala Ser Ala Ser Thr Trp Ser 1520 1525 Asp Gly Gly Ser Val Arg Pro Ser Tyr Trp Leu Val Leu His Asn Leu Thr Pro Gln Ile Asp Gly Ser Thr Leu Arg Thr Ile Cys Met 1555 Gln His Gly Pro Leu Leu Thr Phe His Leu Asn Leu Thr Gln Gly 1570 Thr Ala Leu Ile Arg Tyr Ser Thr Lys Gln Glu Ala Ala Lys Ala 1580 1585 1590 Gln Thr Ala Leu His Met Cys Val Leu Gly Asn Thr Thr Ile Leu 1600 1595 Ala Glu Phe Ala Thr Asp Asp Glu Val Ser Arg Phe Leu Ala Gln 1610 1615 1620 Ala Gln Pro Pro Thr Pro Ala Ala Thr Pro Ser Ala Pro Ala Ala 1630 Gly Trp Gln Ser Leu Glu Thr Gly Gln Asn Gln Ser Asp Pro Val 1645 1640 Gly Pro Ala Leu Asn Leu Phe Gly Gly Ser Thr Gly Leu Gly Gln Trp Ser Ser Ser Ala Gly Gly Ser Ser Gly Ala Asp Leu Ala Gly 1675 1670 Ala Ser Leu Trp Gly Pro Pro Asn Tyr Ser Ser Leu Trp Gly 1690 Val Pro Thr Val Glu Asp Pro His Arg Met Gly Ser Pro Ala Pro 1705 1710 1700 Leu Leu Pro Gly Asp Leu Leu Gly Gly Gly Ser Asp Ser Ile

1720

<210> 12 <211> 3830 <212> PRT

1715

<213> Homo sapiens

<400> 12

Met Ala Phe Val Ala Thr Gln Gly Ala Thr Val Val Asp Gln Thr Thr 1 10 15

Leu Met Lys Lys Tyr Leu Gln Phe Val Ala Ala Leu Thr Asp Val Asn 20 25 30

Thr Pro Asp Glu Thr Lys Leu Lys Met Met Gln Glu Val Ser Glu Asn  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Phe Glu Asn Val Thr Ser Ser Pro Gln Tyr Ser Thr Phe Leu Glu His 50 60

Ile Ile Pro Arg Phe Leu Thr Phe Leu Gln Asp Gly Glu Val Gln Phe 65 70 75 80

Leu Gln Glu Lys Pro Ala Gln Gln Leu Arg Lys Leu Val Leu Glu Ile 85 90 95

Ile His Arg Ile Pro Thr Asn Glu His Leu Arg Pro His Thr Lys Asn 100 105 110

Val Leu Ser Val Met Phe Arg Phe Leu Glu Thr Glu Asn Glu Glu Asn 115 120 125

Val Leu Ile Cys Leu Arg Ile Ile Ile Glu Leu His Lys Gln Phe Arg 130 135 140

Pro Pro Ile Thr Gln Glu Ile His His Phe Leu Asp Phe Val Lys Gln 145 150 155 160

Ile Tyr Lys Glu Leu Pro Lys Val Val Asn Arg Tyr Phe Glu Asn Pro 165 170 175

Gln Val Ile Pro Glu Asn Thr Val Pro Pro Pro Glu Met Val Gly Met 180 185 190

Ile Thr Thr Ile Ala Val Lys Val Asn Pro Glu Arg Glu Asp Ser Glu 195 200 205

Thr Arg Thr His Ser Ile Ile Pro Arg Gly Ser Leu Ser Leu Lys Val 210 215 220

Leu Ala Glu Leu Pro Ile Ile Val Val Leu Met Tyr Gln Leu Tyr Lys 225 230 235 240

Leu Asn Ile His Asn Val Val Ala Glu Phe Val Pro Leu Ile Met Asn 245 250 255

Thr Ile Ala Ile Gln Val Ser Ala Gln Ala Arg Gln His Lys Leu Tyr

Asn Lys Glu Leu Tyr Ala Asp Phe Ile Ala Ala Gln Ile Lys Thr Leu Ser Phe Leu Ala Tyr Ile Ile Arg Ile Tyr Gln Glu Leu Val Thr Lys Tyr Ser Gln Gln Met Val Lys Gly Met Leu Gln Leu Leu Ser Asn Cys 315 Pro Ala Glu Thr Ala His Leu Arg Lys Glu Leu Leu Ile Ala Ala Lys His Ile Leu Thr Thr Glu Leu Arg Asn Gln Phe Ile Pro Cys Met Asp 345 Lys Leu Phe Asp Glu Ser Ile Leu Ile Gly Ser Gly Tyr Thr Ala Arg Glu Thr Leu Arg Pro Leu Ala Tyr Ser Thr Leu Ala Asp Leu Val His 375 His Val Arg Gln His Leu Pro Leu Ser Asp Leu Ser Leu Ala Val Gln Leu Phe Ala Lys Asn Ile Asp Asp Glu Ser Leu Pro Ser Ser Ile Gln 410 Thr Met Ser Cys Lys Leu Leu Leu Asn Leu Val Asp Cys Ile Arg Ser Lys Ser Glu Gln Glu Ser Gly Asn Gly Arg Asp Val Leu Met Arg Met 440 435 Leu Glu Val Phe Val Leu Lys Phe His Thr Ile Ala Arg Tyr Gln Leu 450 Ser Ala Ile Phe Lys Lys Cys Lys Pro Gln Ser Glu Leu Gly Ala Val 465 Glu Ala Ala Leu Pro Gly Val Pro Thr Ala Pro Ala Ala Pro Gly Pro Ala Pro Ser Pro Ala Pro Val Pro Ala Pro Pro Pro Pro Pro Pro Pro 500 505 Pro Pro Pro Ala Thr Pro Val Thr Pro Ala Pro Val Pro Pro Phe Glu Lys Gln Gly Glu Lys Asp Lys Glu Asp Lys Gln Thr Phe Gln Val Thr

535

Asp Cys Arg Ser Leu Val Lys Thr Leu Val Cys Gly Val Lys Thr Ile 545 550 555 560

Thr Trp Gly Ile Thr Ser Cys Lys Ala Pro Gly Glu Ala Gln Phe Ile 565 570 575

Pro Asn Lys Gln Leu Gln Pro Lys Glu Thr Gln Ile Tyr Ile Lys Leu 580 585 590

Val Lys Tyr Ala Met Gln Ala Leu Asp Ile Tyr Gln Val Gln Ile Ala 595 600 605

Gly Asn Gly Gln Thr Tyr Ile Arg Val Ala Asn Cys Gln Thr Val Arg 610 615 620

Met Lys Glu Glu Lys Glu Val Leu Glu His Phe Ala Gly Val Phe Thr 625 630 635 640

Met Met Asn Pro Leu Thr Phe Lys Glu Ile Phe Gln Thr Thr Val Pro 645 650 655

Tyr Met Val Glu Arg Ile Ser Lys Asn Tyr Ala Leu Gln Ile Val Ala 660 665 670

Asn Ser Phe Leu Ala Asn Pro Thr Thr Ser Ala Leu Phe Ala Thr Ile 675 680 685

Leu Val Glu Tyr Leu Leu Asp Arg Leu Pro Glu Met Gly Ser Asn Val 690 695 700

Glu Leu Ser Asn Leu Tyr Leu Lys Leu Phe Lys Leu Val Phe Gly Ser 705 710 715 720

Val Ser Leu Phe Ala Ala Glu As<br/>n Glu Gln Met Leu Lys Pro His Leu 725 730 735

His Lys Ile Val Asn Ser Ser Met Glu Leu Ala Gln Thr Ala Lys Glu 740 745 750

Pro Tyr Asn Tyr Phe Leu Leu Leu Arg Ala Leu Phe Arg Ser Ile Gly 755 760 765

Gly Gly Ser His Asp Leu Leu Tyr Gln Glu Phe Leu Pro 1770 775 780

Asn Leu Leu Gln Gly Leu Asn Met Leu Gln Ser Gly Leu His Lys Gln 785 790 795 800

His Met Lys Asp Leu Phe Val Glu Leu Cys Leu Thr Val Pro Val Arg 805 810 815

Leu Ser Ser Leu Leu Pro Tyr Leu Pro Met Leu Met Asp Pro Leu Val 820 825 830

Ser Ala Leu Asn Gly Ser Gln Thr Leu Val Ser Gln Gly Leu Arg Thr 835 840 845

Leu Glu Leu Cys Val Asp Asn Leu Gln Pro Asp Phe Leu Tyr Asp His 850 855 860

Ile Gln Pro Val Arg Ala Glu Leu Met Gln Ala Leu Trp Arg Thr Leu 865 870 875 880

Arg Asn Pro Ala Asp Ser Ile Ser His Val Ala Tyr Arg Val Leu Gly 885 890 895

Lys Phe Gly Gly Ser Asn Arg Lys Met Leu Lys Glu Ser Gln Lys Leu 900 905 910

His Tyr Val Val Thr Glu Val Gln Gly Pro Ser Ile Thr Val Glu Phe 915 920 925

Ser Asp Cys Lys Ala Ser Leu Gln Leu Pro Met Glu Lys Ala Ile Glu 930 935 940

Thr Ala Leu Asp Cys Leu Lys Ser Ala Asn Thr Glu Pro Tyr Tyr Arg 945 950 955 960

Arg Gln Ala Trp Glu Val Ile Lys Cys Phe Leu Val Ala Met Met Ser 965 970 975

Leu Glu Asp Asn Lys His Ala Leu Tyr Gln Leu Leu Ala His Pro Asn 980 985 990

Phe Thr Glu Lys Thr Ile Pro Asn Val Ile Ile Ser His Arg Tyr Lys 995 1000 1005

Ala Gln Asp Thr Pro Ala Arg Lys Thr Phe Glu Gln Ala Leu Thr 1010 \$1015\$

Leu Pro Phe Val Ala Ser Leu Ile Arg His Tyr Thr Met Val Ala 1040 1045 1050

Val Ala Gln Gln Cys Gly Pro Phe Leu Leu Pro Cys Tyr Gln Val 1055 1060 1065

Gly Sér Gln Pro Ser Thr Ala Met Phe His Ser Glu Glu Asn Gly 1070 1080 Ser Lys Gly Met Asp Pro Leu Val Leu Ile Asp Ala Ile Ala Ile 1085 1090 1095

e.i

Cys Met Ala Tyr Glu Glu Lys Glu Leu Cys Lys Ile Gly Glu Val 1100 1105 1110

Ala Leu Ala Val Ile Phe Asp Val Ala Ser Ile Ile Leu Gly Ser 1115 1120 1125

Lys Glu Arg Ala Cys Gln Leu Pro Leu Phe Ser Tyr Ile Val Glu 1130 1135 1140

Arg Leu Cys Ala Cys Cys Tyr Glu Gl<br/>n Ala Trp Tyr Ala Lys Leu 1145 1150 1155

Gly Gly Val Val Ser Ile Lys Phe Leu Met Glu Arg Leu Pro Leu 1160 1165 1170

Thr Trp Val Leu Gln Asn Gln Gln Thr Phe Leu Lys Ala Leu Leu 1175 1180 1185

Phe Val Met Met Asp Leu Thr Gly Glu Val Ser Asn Gly Ala Val 1190 1195 1200

Ala Met Ala Lys Thr Thr Leu Glu Gln Leu Leu Met Arg Cys Ala 1205 1210 1215

Thr Pro Leu Lys Asp Glu Glu Arg Ala Glu Glu Ile Val Ala Ala 1220 1230

Gln Glu Lys Ser Phe His His Val Thr His Asp Leu Val Arg Glu 1235 1240 1245

Val Thr Ser Pro Asn Ser Thr Val Arg Lys Gln Ala Met His Ser 1250 1255 1260

Leu Gln Val Leu Ala Gln Val Thr Gly Lys Ser Val Thr Val Ile 1265 1270 1275

Met Glu Pro His Lys Glu Val Leu Gln Asp Met Val Pro Pro Lys 1280 1285 1290

Lys His Leu Leu Arg His Gln Pro Ala Asn Ala Gln Ile Gly Leu 1295 1300 1305

Met Glu Gly Asn Thr Phe Cys Thr Thr Leu Gln Pro Arg Leu Phe 1310 1320

Thr Met Asp Leu Asn Val Val Glu His Lys Val Phe Tyr Thr Glu 1325 1330 1335

Leu Leu Asn Leu Cys Glu Ala Glu Asp Ser Ala Leu Thr Lys Leu

1340 1345 1350

I	Pro	Cys 1355	Tyr	Lys	Ser	Leu	Pro 1360	Ser	Leu	Val	Pro	Leu 1365	Arg	Ile	Ala
1	Ala	Leu 1370	Asn	Ala	Leu	Ala	Ala 1375	Cys	Asn	Tyr	Leu	Pro 1380	Gln	Ser	Arg
(	Glu	Lys 1385		Ile	Ala	Ala	Leu 1390		Lys	Ala	Leu	Asn 1395	Ser	Thr	Asn
5	Ser	Glu 1400	Leu	Gln	Glu	Ala	Gly 1405	Glu	Ala	Cys	Met	Arg 1410	Lys	Phe	Leu
(	Glu	Gly 1415		Thr	Ile	Glu	Val 1420		Gln	Ile	His	Thr 1425		Met	Arg
1	Pro	Leu 1430		Met	Met	Leu	Gly 1435		Tyr	Arg	Ser	Leu 1440	Thr	Leu	Asn
,	Val	Val 1445	Asn	Arg	Leu	Thr	Ser 1450	Val	Thr	Arg	Leu	Phe 1455	Pro	Asn	Ser
]	Phe	Asn 1460	_	Lys	Phe	Cys	Asp 1465		Met	Met	Gln	His 1470	Leu	Arg	Lys
	Гrр	Met 1475		Val	Val	Val	Ile 1480	Thr	His	Lys	Gly	Gly 1485	Gln	Arg	Ser
ì	Asp	Gly 1490		Glu	Met	Lys	Ile 1495		Ser	Ala	Ile	Ile 1500	Asn	Leu	Phe
J	His	Leu 1505	Ile	Pro	Ala	Ala	Pro 1510	Gln	Thr	Leu	Val	Lys 1515	Pro	Leu	Leu
(	Glu	Val 1520	Val	Met	Lys	Thr	Glu 1525	Arg	Ala	Met	Leu	Ile 1530	Glu	Ala	Gly
:	Ser	Pro 1535	Phe	Arg	Glu	Pro	Leu 1540	Ile	Lys	Phe	Leu	Thr 1545	Arg	His	Pro
:	Ser	Gln 1550	Thr	Val	Glu	Leu	Phe 1555	Met	Met	Glu	Ala	Thr 1560	Leu	Asn	Asp
	Pro	Gln 1565	Trp	Ser	Arg	Met	Phe 1570	Met	Ser	Phe	Leu	Lys 1575		Lys	Asp
i	Ala	Arg 1580	Pro	Leu	Arg	Asp	Val 1585	Leu	Ala	Ala	Asn	Pro 1590	Asn	Arg	Phe
	Ile	Thr 1595	Leu	Leu	Leu	Pro	Gly 1600	Gly	Ala	Gln	Thr	Ala 1605	Val	Arg	Pro

Gly Ser Pro Ser Thr Ser Thr Met Arg Leu Asp Leu Gln Phe Gln 1610 1620

Ala Ile Lys Ile Ile Ser Ile Ile Val Lys Asn Asp Asp Ser Trp 1625 1630 1635

Leu Ala Ser Gln His Ser Leu Val Ser Gln Leu Arg Arg Val Trp 1640 1645 1650

Val Ser Glu Asn Phe Gln Glu Arg His Arg Lys Glu Asn Met Ala 1655 1660 1665

Ala Thr Asn Trp Lys Glu Pro Lys Leu Leu Ala Tyr Cys Leu Leu 1670 1680

Asn Tyr Cys Lys Arg Asn Tyr Gly Asp Ile Glu Leu Leu Phe Gln 1685 1690 1695

Leu Leu Arg Ala Phe Thr Gly Arg Phe Leu Cys Asn Met Thr Phe 1700 1710 1710

Leu Lys Glu Tyr Met Glu Glu Glu Ile Pro Lys Asn Tyr Ser Ile 1715 1720 1725

Ala Gln Lys Arg Ala Leu Phe Phe Arg Phe Val Asp Phe Asn Asp 1730 1735 1740

Pro Asn Phe Gly Asp Glu Leu Lys Ala Lys Val Leu Gln His Ile 1745 \$1750\$

Leu Asn Pro Ala Phe Leu Tyr Ser Phe Glu Lys Gly Glu Gly Glu 1760 1765 1770

Gln Leu Leu Gly Pro Pro Asn Pro Glu Gly Asp Asn Pro Glu Ser 1775 1780 1785

Ile Thr Ser Val Phe Ile Thr Lys Val Leu Asp Pro Glu Lys Gln 1790 1795 1800

Ala Asp Met Leu Asp Ser Leu Arg Ile Tyr Leu Leu Gln Tyr Ala 1805 1810 1815

Thr Leu Leu Val Glu His Ala Pro His His Ile His Asp Asn Asn 1820 1825 1830

Lys Asn Arg Asn Ser Lys Leu Arg Arg Leu Met Thr Phe Ala Trp 1835 1840 1845

Pro Cys Leu Leu Ser Lys Ala Cys Val Asp Pro Ala Cys Lys Tyr 1850 1855 1860 Ser Gly His Leu Leu Leu Ala His Ile Ile Ala Lys Phe Ala Ile 1865 1870 1875

His Lys Lys Ile Val Leu Gln Val Phe His Ser Leu Leu Lys Ala 1880 1885 1890

His Ala Met Glu Ala Arg Ala Ile Val Arg Gln Ala Met Ala Ile 1895 1900 1905

Leu Thr Pro Ala Val Pro Ala Arg Met Glu Asp Gly His Gln Met 1910 . 1915 1920

Leu Thr His Trp Thr Arg Lys Ile Ile Val Glu Glu Gly His Thr 1925 1930 1935

Val Pro Gln Leu Val His Ile Leu His Leu Ile Val Gln His Phe 1940 1945 1950

Lys Val Tyr Tyr Pro Val Arg His His Leu Val Gln His Met Val 1955 1960 1965

Ser Ala Met Gln Arg Leu Gly Phe Thr Pro Ser Val Thr Ile Glu 1970 1975 1980

Gln Arg Arg Leu Ala Val Asp Leu Ser Glu Val Val Ile Lys Trp 1985 1990 1995

Glu Leu Gln Arg Ile Lys Asp Gln Gln Pro Asp Ser Asp Met Asp 2000 2010

Pro Asn Ser Ser Gly Glu Gly Val Asn Ser Val Ser Ser Ser Ile 2015 2020 2025

Lys Arg Gly Leu Ser Val Asp Ser Ala Gln Glu Val Lys Arg Phe 2030 2035 2040

Arg Thr Ala Thr Gly Ala Ile Ser Ala Val Phe Gly Arg Ser Gln 2045 2050 2055

Ser Leu Pro Gly Ala Asp Ser Leu Leu Ala Lys Pro Ile Asp Lys 2060 2065 2070

Gln His Thr Asp Thr Val Val Asn Phe Leu Ile Arg Val Ala Cys 2075 2080 2085

Gln Val Asn Asp Asn Thr Asn Thr Ala Gly Ser Pro Gly Glu Val 2090 2095 2100

Leu Ser Arg Arg Cys Val Asn Leu Leu Lys Thr Ala Leu Arg Pro 2105 2110 2115 Asp Met Trp Pro Lys Ser Glu Leu Lys Leu Gln Trp Phe Asp Lys 2120 Met Thr Val Glu Gln Pro Asn Gln Val Asn Tyr Gly Asn 2135

Ile Cys Thr Gly Leu Glu Val Leu Ser Phe Leu Leu Thr Val Leu 2150 2155 2160

Gln Ser Pro Ala Ile Leu Ser Ser Phe Lys Pro Leu Gln Arg Gly 2165 2170 2175

Ile Ala Ala Cys Met Thr Cys Gly Asn Thr Lys Val Leu Arg Ala 2180 2185 2190

Val His Ser Leu Leu Ser Arg Leu Met Ser Ile Phe Pro Thr Glu 2195 2200 2205

Pro Ser Thr Ser Ser Val Ala Ser Lys Tyr Glu Glu Leu Glu Cys 2210 2215 2220

Leu Tyr Ala Ala Val Gly Lys Val Ile Tyr Glu Gly Leu Thr Asn 2225 2230 2235

Tyr Glu Lys Ala Thr Asn Ala Asn Pro Ser Gln Leu Phe Gly Thr 2240  $\phantom{0}2245$   $\phantom{0}2250$ 

Leu Met Ile Leu Lys Ser Ala Cys Ser Asn Asn Pro Ser Tyr Ile 2255 2260 2265

Asp Arg Leu Ile Ser Val Phe Met Arg Ser Leu Gln Lys Met Val 2270 2280

Arg Glu His Leu Asn Pro Gln Ala Ala Ser Gly Ser Thr Glu Ala 2285 2290 2295

Thr Ser Gly Thr Ser Glu Leu Val Met Leu Ser Leu Glu Leu Val 2300 2305 2310

Lys Thr  $\mbox{Arg}$  Leu Ala Val Met Ser Met Glu Met Arg  $\mbox{Lys}$  Asn Phe 2315 2320 2325

Ile Gln Ala Ile Leu Thr Ser Leu Ile Glu Lys Ser Pro Asp Ala 2330 2340

Lys Ile Leu Arg Ala Val Val Lys Ile Val Glu Glu Trp Val Lys 2345 2350 2355

Asn Asn Ser Pro Met Ala Ala Asn Gln Thr Pro Thr Leu Arg Glu 2360 2365 2370

Lys Ser Ile Leu Leu Val Lys Met Met Thr Tyr Ile Glu Lys Arg

2375 2380 2385

Phe	Pro 2390	Glu	Asp	Leu	Glu	Leu 2395	Asn	Ala	Gln	Phe	Leu 2400	Asp	Leu	Val
Asn	Tyr 2405	Val	Tyr	Arg		Glu 2410	Thr	Leu	Ser	Gly	Ser 2415	Glu	Leu	Thr
Ala	Lys 2420	Leu	Glu	Pro		Phe 2425	Leu	Ser	Gly	Leu	Arg 2430	Суѕ	Ala	Gln
Pro	Leu 2435	Ile	Arg	Ala	Lys	Phe 2440	Phe	Glu	Val	Phe	Asp 2445	Asn	Ser	Met
Lys	Arg 2450	Arg	Val	Tyr	Glu	Arg 2455	Leu	Leu	Tyr	Val	Thr 2460	Cys	Ser	Gln
Asn	Trp 2465	Glu	Ala	Met	Gly	Asn 2470	His	Phe	Trp	Ile	Lys 2475	Gln	Cys	Ile
Glu	Leu 2480		Leu	Ala	Val	Cys 2485	Glu	Lys	Ser	Thr	Pro 2490	Ile	Gly	Thr
Ser	Cys 2495	Gln	Gly	Ala	Met	Leu 2500	Pro	Ser	Ile	Thr	Asn 2505	Val	Ile	Asn
Leu	Ala 2510		Ser	His	Asp	Arg 2515	Ala	Ala	Phe	Ala	Met 2520		Thr	His
Val	Lys 2525		Glu	Pro	Arg	Glu 2530	Arg	Glu	Asn	Ser	Glu 2535	Ser	Lys	Glu
Glu	Asp 2540	Val	Glu	Ile	Asp	Ile 2545	Glu	Leu	Ala	Pro	Gly 2550	Asp	Gln	Thr
Ser	Thr 2555	Pro	Lys	Thr	Lys	Glu 2560	Leu	Ser	Glu	Lys	Asp 2565	Ile	Gly	Asn
Gln	Leu 2570	His	Met	Leu	Thr	Asn 2575	Arg	His	Asp	Lys	Phe 2580	Leu	Asp	Thr
Leu	Arg 2585	Glu	Val	Lys	Thr	Gly 2590	Ala	Leu	Leu	Ser	Ala 2595	Phe	Val	Gln
Leu	Cys 2600		Ile	Ser	Thr	Thr 2605	Leu	Ala	Glu	Lys	Thr 2610	Trp	Val	Gln
Leu	Phe 2615	Pro	Arg	Leu	Trp	Lys 2620	Ile	Leu	Ser	Asp	Arg 2625	Gln	Gln	His
Ala	Leu 2630	Ala	Gly	Glu	Ile	Ser 2635	Pro	Phe	Leu	Cys	Ser 2640	Gly	Ser	His

Gln Val Gln Arg Asp Cys Gln Pro Ser Ala Leu Asn Cys Phe Val 2645 2650 2655

Glu Ala Met Ser Gln Cys Val Pro Pro Ile Pro Ile Arg Pro Cys 2660 2665 2670

Val Leu Lys Tyr Leu Gly Lys Thr His Asn Leu Trp Phe Arg Ser 2675 2680 2685

Thr Leu Met Leu Glu His Gln Ala Phe Glu Lys Gly Leu Ser Leu 2690 2695 2700

Gln Ile Lys Pro Lys Gln Thr Thr Glu Phe Tyr Glu Gln Glu Ser 2705 2710 2715

Ile Thr Pro Pro Gln Gln Glu Ile Leu Asp Ser Leu Ala Glu Leu 2720 2725 2730

Tyr Ser Leu Leu Gln Glu Glu Asp Met Trp Ala Gly Leu Trp Gln 2735 2740 2745

Lys Arg Cys Lys Tyr Ser Glu Thr Ala Thr Ala Ile Ala Tyr Glu 2750  $\phantom{000}2755$   $\phantom{0000}2760$ 

Gln His Gly Phe Phe Glu Gln Ala Gln Glu Ser Tyr Glu Lys Ala 2765 2770 2775

Met Asp Lys Ala Lys Lys Glu His Glu Arg Ser Asn Ala Ser Pro 2780 2785 2790

Ala Ile Phe Pro Glu Tyr Gln Leu Trp Glu Asp His Trp Ile Arg 2795 2800 2805

Cys Ser Lys Glu Leu Asn Gln Trp Glu Ala Leu Thr Glu Tyr Gly 2810 2820

Gln Ser Lys Gly His Ile Asn Pro Tyr Leu Val Leu Glu Cys Ala 2825 2830 2835

Trp Arg Val Ser Asn Trp Thr Ala Met Lys Glu Ala Leu Val Gln 2840 2845 2850

Val Glu Val Ser Cys Pro Lys Glu Met Ala Trp Lys Val Asn Met 2855 2860 2865

Tyr Arg Gly Tyr Leu Ala Ile Cys His Pro Glu Glu Gln Gln Leu 2870 2880

Ser Phe Ile Glu Arg Leu Val Glu Met Ala Ser Ser Leu Ala Ile 2885 2890 2895 Arg Glu Trp Arg Arg Leu Pro His Val Val Ser His Val His Thr 2900 2905 2910

Pro Leu Leu Gln Ala Ala Gln Gln Ile Ile Glu Leu Gln Glu Ala 2915 2920 2925

Ala Gln Ile Asn Ala Gly Leu Gln Pro Thr Asn Leu Gly Arg Asn 2930 2940

Asn Ser Leu His Asp Met Lys Thr Val Val Lys Thr Trp Arg Asn 2945 2950 2955

Arg Leu Pro Ile Val Ser Asp Asp Leu Ser His Trp Ser Ser Ile 2960 2965 2970

Phe Met Trp Arg Gln His His Tyr Gln Ala Ile Val Thr Ala Tyr 2975 2980 2985

Glu Asn Ser Ser Gln His Asp Pro Ser Ser Asn Asn Ala Met Leu 2990 2995 3000

Gly Val His Ala Ser Ala Ser Ala Ile Ile Gln Tyr Gly Lys Ile 3005 3010 3015

Ala Arg Lys Gln Gly Leu Val Asn Val Ala Leu Asp Ile Leu Ser 3020 3025 3030

Lys Ile Arg Gln Gln Val Lys Cys Tyr Leu Gln Leu Ala Gly Val 3050 3060

Thr Asn Leu Lys Tyr Phe Thr Lys Glu Met Thr Ala Glu Phe Tyr 3080 3085 3090

Ala Leu Lys Gly Met Phe Leu Ala Gln Ile Asn Lys Ser Glu Glu 3095 3100 3105

Ala Asn Lys Ala Phe Ser Ala Ala Val Gln Met His Asp Val Leu 3110 3115 3120

Val Lys Ala Trp Ala Met Trp Gly Asp Tyr Leu Glu Asn Ile Phe 3125 3130 3135

Val Lys Glu Arg Gln Leu His Leu Gly Val Ser Ala Ile Thr Cys 3140 3145 3150 Tyr Leu His Ala Cys Arg His Gln Asn Glu Ser Lys Ser Arg Lys 3155 3160 3165

Tyr Leu Ala Lys Val Leu Trp Leu Leu Ser Phe Asp Asp Asp Lys 3170 3180

Asn Thr Leu Ala Asp Ala Val Asp Lys Tyr Cys Ile Gly Val Pro 3185 3190 3195

Pro Ile Gln Trp Leu Ala Trp Ile Pro Gln Leu Leu Thr Cys Leu 3200 3205 3210

Val Gly Ser Glu Gly Lys Leu Leu Asn Leu Ile Ser Gln Val 3215 3220 3225

Gly Arg Val Tyr Pro Gln Ala Val Tyr Phe Pro Ile Arg Thr Leu 3230 3240

Tyr Leu Thr Leu Lys Ile Glu Gln Arg Glu Arg Tyr Lys Ser Asp 3245 3250 3255

Pro Gly Pro Ile Arg Ala Thr Ala Pro Met Trp Arg Cys Ser Arg 3260 3265 3270

Ile Met His Met Gln Arg Glu Leu His Pro Thr Leu Leu Ser Ser 3275 3280 3285

Leu Glu Gly Ile Val Asp Gln Met Val Trp Phe Arg Glu Asn Trp 3290 3295 3300

His Glu Glu Val Leu Arg Gln Leu Gln Gln Gly Leu Ala Lys Cys 3305 3310 3315

Tyr Ser Val Ala Phe Glu Lys Ser Gly Ala Val Ser Asp Ala Lys 3320 3335 3330

Ile Thr Pro His Thr Leu Asn Phe Val Lys Lys Leu Val Ser Thr 3335 3340 3345

Phe Gly Val Gly Leu Glu Asn Val Ser Asn Val Ser Thr Met Phe 3350 3360

Ser Ser Ala Ala Ser Glu Ser Leu Ala Arg Arg Ala Gln Ala Thr 3365 3370 3375

Ala Gln Asp Pro Val Phe Gln Lys Leu Lys Gly Gln Phe Thr Thr 3380 3385 3390

Asp Phe Asp Phe Ser Val Pro Gly Ser Met Lys Leu His Asn Leu 3395 3400 3405

Ile Ser Lys Leu Lys Lys Trp Ile Lys Ile Leu Glu Ala Lys Thr

3410 3415 3420

Lys	Gln 3425	Leu	Pro	Lys	Phe	Phe 3430	Leu	Ile	Glu	Glu	Lys 3435	Cys	Arg	Phe
Leu	Ser 3440	Asn	Phe	Ser	Ala	Gln 3445	Thr	Ala	Glu	Val	Glu 3450	Ile	Pro	Gly
Glu	Phe 3455	Leu	Met	Pro	Lys	Pro 3460	Thr	His	Tyr	Tyr	Ile 3465	Lys	Ile	Ala
Arg	Phe 3470	Met	Pro	Arg	Val	Glu 3475	Ile	Val	Gln	Lys	His 3480	Asn	Thr	Ala
Ala	Arg 3485	Arg	Leu	Tyr	Ile	Arg 3490	Gly	His	Asn	Gly	Lys 3495		Tyr	Pro
Tyr	Leu 3500	Val	Met	Asn	Asp	Ala 3505	Cys	Leu	Thr	Glu	Ser 3510	Arg	Arg	Glu
Glu	Arg 3515		Leu	Gln	Leu	Leu 3520	Arg	Leu	Leu	Asn	Pro 3525		Leu	Glu
Lys	Arg 3530		Glu	Thr	Thr	Lys 3535	Arg	His	Leu	Phe	Phe 3540	Thr	Val	Pro
Arg	Val 3545	Val	Ala	Val	Ser	Pro 3550	Gln	Met	Arg	Leu	Val 3555	Glu	Asp	Asn
Pro	Ser 3560		Leu	Ser	Leu	Val 3565	Glu	Ile	Tyr	Lys	Gln 3570	Arg	Cys	Ala
Lys	Lys 3575	Gly	Ile	Glu	His	Asp 3580	Asn	Pro	Ile	Ser	Arg 3585	Tyr	Tyr	Asp
Arg	Leu 3590		Thr	Val	Gln	Ala 3595	Arg	Gly	Thr	Gln	Ala 3600		His	Gln
Val	Leu 3605	Arg	Asp	Ile	Leu	Lys 3610	Glu	Val	Gln	Ser	Asn 3615	Met	Val	Pro
Arg	Ser 3620	Met	Leu	Lys	Glu	Trp 3625	Ala	Leu	His	Thr	Phe 3630	Pro	Asn	Ala
Thr	Asp 3635		Trp	Thr	Phe	Arg 3640		Met	Phe	Thr	Ile 3645		Leu	Ala
Leu	Ile 3650	_	Phe	Ala	Glu	Phe 3655	Val	Leu	His	Leu	Asn 3660		Leu	Asn
Pro	Glu 3665		Leu	Gln	Ile	Ala 3670		Asp	Thr	Gly	Lys 3675		Asn	Val

Ala Tyr Phe Arg Phe Asp Ile Asn Asp Ala Thr Gly Asp Leu Asp 3680 3685 3690

Ala Asn Arg Pro Val Pro Phe Arg Leu Thr Pro Asn Ile Ser Glu 3695 3700 3705

Phe Leu Thr Thr Ile Gly Val Ser Gly Pro Leu Thr Ala Ser Met 3710 3715 3720

Ile Ala Val Ala Arg Cys Phe Ala Gln Pro Asn Phe Lys Val Asp 3725 3730 3735

Gly Ile Leu Lys Thr Val Leu Arg Asp Glu Ile Ile Ala Trp His 3740 3745 3750

Lys Lys Thr Gln Glu Asp Thr Ser Ser Pro Leu Ser Ala Ala Gly 3755 3760 3765

Gln Pro Glu Asn Met Asp Ser Gln Gln Leu Val Ser Leu Val Gln 3770  $\phantom{\bigg|}3775\phantom{\bigg|}3780\phantom{\bigg|}$ 

Lys Ala Val Thr Ala Ile Met Thr Arg Leu His Asn Leu Ala Gln 3785 3790 3795

Phe Glu Gly Gly Glu Ser Lys Val Asn Thr Leu Val Ala Ala Ala 3800 \$3805

Trp Leu 3830

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<213> Homo sapiens

<400> 13

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Val Met Lys Gln Phe Ala Phe Val His Met Arg Glu Asn Ala Gly Ala 35 40 45

Leu Arg Ala Ile Glu Ala Leu His Gly His Glu Leu Arg Pro Gly Arg 50 55 60

Ala Leu Val Val Glu Met Ser Arg Pro Arg Pro Leu Asn Thr Trp Lys 75 75 80

Ile Phe Val Gly Asn Val Ser Ala Ala Cys Thr Ser Gln Glu Leu Arg

Ser Leu Phe Glu Arg Arg Gly Arg Val Ile Glu Cys Asp Val Val Lys

Asp Tyr Ala Phe Val His Met Glu Lys Glu Ala Asp Ala Lys Ala Ala 115 120 125

Ile Ala Gln Leu Asn Gly Lys Glu Val Lys Gly Lys Arg Ile Asn Val 130 135 140

Glu Leu Ser Thr Lys Gly Gln Lys Lys Gly Pro Gly Leu Ala Val Gln 145 150 155 160

Ser Gly Asp Lys Thr Lys Lys Pro Gly Ala Gly Asp Thr Ala Phe Pro 165 170 175

Gly Thr Gly Gly Phe Ser Ala Thr Phe Asp Tyr Gln Gln Ala Phe Gly 180 185 190

Asn Ser Thr Gly Gly Phe Asp Gly Gln Ala Arg Gln Pro Thr Pro Pro 195 200 205

Phe Phe Gly Arg Asp Arg Ser Pro Leu Arg Arg Ser Pro Pro Arg Ala 210 215 220

Ser Tyr Val Ala Pro Leu Thr Ala Gln Pro Ala Thr Tyr Arg Ala Gln 225 230 235 240

Pro Ser Val Ser Leu Gly Ala Ala Tyr Arg Ala Gln Pro Ser Ala Ser 245 250 255

Leu Gly Val Gly Tyr Arg Thr Gln Pro Met Thr Ala Gln Ala Ala Ser 260 265 270

Tyr Arg Ala Gln Pro Ser Val Ser Leu Gly Ala Pro Tyr Arg Gly Gln 275 280 285

Leu Ala Ser Pro Ser Ser Gln Ser Ala Ala Ala Ser Ser Leu Gly Pro 290 295 300

Tyr Gly Gly Ala Gln Pro Ser Ala Ser Ala Leu Ser Ser Tyr Gly Gly 305 310 315 320

Gln Ala Ala Ala Ser Ser Leu Asn Ser Tyr Gly Ala Gln Gly Ser 325 330 335

Ser Leu Ala Ser Tyr Gly Asn Gln Pro Ser Ser Tyr Gly Ala Gln Ala

Ala Ser Ser Tyr Gly Val Arg Ala Ala Ala Ser Ser Tyr Asn Thr Gln Gly Ala Ala Ser Ser Leu Gly Ser Tyr Gly Ala Gln Ala Ala Ser Tyr Gly Ala Gln Ser Ala Ala Ser Ser Leu Ala Tyr Gly Ala Gln Ala Ala 395 390 385 Ser Tyr Asn Ala Gln Pro Ser Ala Ser Tyr Asn Ala Gln Ser Ala Pro Tyr Ala Ala Gln Gln Ala Ala Ser Tyr Ser Ser Gln Pro Ala Ala Tyr 420 Val Ala Gln Pro Ala Thr Ala Ala Ala Tyr Ala Ser Gln Pro Ala Ala Tyr Ala Ala Gln Ala Thr Thr Pro Met Ala Gly Ser Tyr Gly Ala Gln 455 Pro Val Val Gln Thr Gln Leu Asn Ser Tyr Gly Ala Gln Ala Ser Met Gly Leu Ser Gly Ser Tyr Gly Ala Gln Ser Ala Ala Ala Ala Thr Gly Ser Tyr Gly Ala Ala Ala Ala Tyr Gly Ala Gln Pro Ser Ala Thr Leu 505 Ala Ala Pro Tyr Arg Thr Gln Ser Ser Ala Ser Leu Ala Ala Ser Tyr 525 515 520 Ala Ala Gln Gln His Pro Gln Ala Ala Ala Ser Tyr Arg Gly Gln Pro 530 Gly Asn Ala Tyr Asp Gly Ala Gly Gln Pro Ser Ala Ala Tyr Leu Ser Met Ser Gln Gly Ala Val Ala Asn Ala Asn Ser Thr Pro Pro Pro Tyr Glu Arg Thr Arg Leu Ser Pro Pro Arg Ala Ser Tyr Asp Asp Pro Tyr 585 580 Lys Lys Ala Val Ala Met Ser Lys Arg Tyr Gly Ser Asp Arg Arg Leu Ala Glu Leu Ser Asp Tyr Arg Arg Leu Ser Glu Ser Gln Leu Ser Phe

615

Arg Arg Ser Pro Thr Lys Ser Ser Leu Asp Tyr Arg Arg Leu Pro Asp 625 630 635 640

Ala His Ser Asp Tyr Ala Arg Tyr Ser Gly Ser Tyr Asn Asp Tyr Leu 645 650 655

Arg Ala Ala Gln Met His Ser Gly Tyr Gln Arg Arg Met 660 665

<210> 14

<211> 456

<212> PRT

<213> Homo sapiens

<400> 14

Met Lys Ile Glu Glu Val Lys Ser Thr Thr Lys Thr Gln Arg Ile Ala 1 5 10 15

Ser His Ser His Val Lys Gly Leu Gly Leu Asp Glu Ser Gly Leu Ala 20 25 30

Lys Gln Ala Ala Ser Gly Leu Val Gly Gln Glu Asn Ala Arg Glu Ala 35 40 45

Cys Gly Val Ile Val Glu Leu Ile Lys Ser Lys Lys Met Ala Gly Arg 50 55 .60

Ala Val Leu Leu Ala Gly Pro Pro Gly Thr Gly Lys Thr Ala Leu Ala 65 70 75 80

Leu Ala Ile Ala Gl<br/>n Glu Leu Gly Ser Lys Val Pro Phe Cys Pro Met 85 90 95

Val Gly Ser Glu Val Tyr Ser Thr Glu Ile Lys Lys Thr Glu Val Leu 100 105 110

Met Glu Asn Phe Arg Arg Ala Ile Gly Leu Arg Ile Lys Glu Thr Lys 115 120 125

Glu Val Tyr Glu Gly Glu Val Thr Glu Leu Thr Pro Cys Glu Thr Glu 130 135 140

Asn Pro Met Gly Gly Tyr Gly Lys Thr Ile Ser His Val Ile Ile Gly 145 150 155

Leu Lys Thr Ala Lys Gly Thr Lys Gln Leu Lys Leu Asp Pro Ser Ile 165 170 175

Phe Glu Ser Leu Gln Lys Glu Arg Val Glu Ala Gly Asp Val Ile Tyr 180 185 190 Ile Glu Ala Asn Ser Gly Ala Val Lys Arg Gln Gly Arg Cys Asp Thr 195 200 205

Tyr Ala Thr Glu Phe Asp Leu Glu Ala Glu Glu Tyr Val Pro Leu Pro 210 215 220

Lys Gly Asp Val His Lys Lys Lys Glu Ile Ile Gln Asp Val Thr Leu 225 230 235 240

His Asp Leu Asp Val Ala Asn Ala Arg Pro Gln Gly Gly Gln Asp Ile 245 250 255

Leu Ser Met Met Gly Gln Leu Met Lys Pro Lys Lys Thr Glu Ile Thr 260 265 270

Asp Lys Leu Arg Gly Glu Ile Asn Lys Val Val Asn Lys Tyr Ile Asp 275 280 285

Gln Gly Ile Ala Glu Leu Val Pro Gly Val Leu Phe Val Asp Glu Val 290 295 300

His Met Leu Asp Ile Glu Cys Phe Thr Tyr Leu His Arg Ala Leu Glu 305 310 315 320

Ser Ser Ile Ala Pro Ile Val Ile Phe Ala Ser Asn Arg'Gly Asn Cys 325 330 335

Val Ile Arg Gly Thr Glu Asp Ile Thr Ser Pro His Gly Ile Pro Leu 340 345 350

Asp Leu Leu Asp Arg Val Met Ile Ile Arg Thr Met Leu Tyr Thr Pro 355 360 365

Gln Glu Met Lys Gln Ile Ile Lys Ile Arg Ala Gln Thr Glu Gly Ile 370 375 380

Asn Ile Ser Glu Glu Ala Leu Asn His Leu Gly Glu Ile Gly Thr Lys 385 390 395 400

Thr Thr Leu Arg Tyr Ser Val Gln Leu Leu Thr Pro Ala Asn Leu Leu 405 410 415

Ala Lys Ile Asn Gly Lys Asp Ser Ile Glu Lys Glu His Val Glu Glu 420 , 425 430

Ile Ser Glu Leu Phe Tyr Asp Ala Lys Ser Ser Ala Lys Ile Leu Ala 435 440

Asp Gln Gln Asp Lys Tyr Met Lys 450 455 <211> 475

<212> PRT <213> Homo sapiens

<400> 15

Met Leu Pro Gly Pro Ala Leu Arg Gly Pro Gly Pro Ala Gln Tyr Gln

Arg Pro Gly Met Ser Pro Gly Asn Arg Met Pro Met Ala Arg Leu Ala

Gly Gly Thr Pro Cys Trp Leu Pro Ile Trp Cys Ser Ser Ser Ala Ser

Thr Trp His Pro Thr His His Asp Gly Ser Ile Pro Lys Thr Pro Ala

Cys Ala Pro Ala Gln Pro Pro Met Pro Ala Gln Arg Arg Gly Leu Lys

Arg Arg Lys Met Ala Asp Lys Val Leu Pro Gln Arg Ile Arg Glu Leu 85

Val Pro Glu Ser Gln Ala Tyr Met Asp Leu Leu Ala Phe Glu Arg Lys

Leu Asp Gln Thr Ile Ala Arg Lys Arg Met Glu Ile Gln Glu Ala Ile 115

Lys Lys Pro Leu Thr Gln Lys Arg Lys Leu Arg Ile Tyr Ile Ser Asn

Thr Phe Ser Pro Ser Lys Ala Glu Gly Asp Ser Ala Gly Thr Ala Gly 145 155

Thr Pro Gly Gly Thr Pro Ala Gly Asp Lys Val Ala Ser Trp Glu Leu

Arg Val Glu Gly Lys Leu Leu Asp Asp Pro Ser Lys Gln Lys Arg Lys

Phe Ser Ser Phe Phe Lys Ser Leu Val Ile Glu Leu Asp Lys Glu Leu

Tyr Gly Pro Asp Gly His Leu Val Glu Trp Tyr Trp Met Pro Thr Thr 210 215 220

Gln Glu Thr Asp Gly Phe Gln Val Lys Arg Pro Gly Asp Leu Asn Val 225

Lys Cys Thr Leu Leu Met Leu Asp His Gln Pro Pro Gln Tyr Lys 245 250

Leu Asp Pro Arg Leu Ala Arg Leu Leu Gly Val His Thr Gln Thr Arg

Ala Ala Ile Met Gln Ala Leu Trp Leu Tyr Ile Lys His Asn Gln Leu 275 280 285

Gln Asp Gly His Glu Arg Glu Tyr Ile Asn Cys Asn Arg Tyr Phe Arg 290 295 300

Gln Ile Phe Ser Cys Gly Arg Leu Arg Phe Ser Glu Ile Pro Met Lys 305 310 315 320

Leu Ala Gly Leu Leu Gln His Pro Asp Pro Ile Val Ile Asn His Val 325 330 335

Ile Ser Val Asp Pro Asn Asp Gln Lys Lys Thr Ala Cys Tyr Asp Ile 340 345 350

Asp Val Glu Val Asp Asp Pro Leu Lys Ala Gln Met Ser Asn Phe Leu 355 360 365

Ala Ser Thr Thr Asn Gln Gln Glu Ile Ala Ser Leu Asp Val Lys Ile 370 375 380

His Glu Thr Ile Glu Ser Ile Asn Gln Leu Lys Thr Gln Arg Asp Phe 385 390 395 400

Met Leu Ser Phe Ser Thr Asp Pro Gln Asp Phe Ile Gln Glu Trp Leu 405 410 415

Arg Ser Gln Arg Arg Asp Leu Lys Ile Ile Thr Asp Val Thr Gly Asn  $420 \hspace{1.5cm} 425 \hspace{1.5cm} 430$ 

Pro Glu Glu Glu Arg Arg Ala Ala Phe Tyr His Gln Pro Trp Ala Gln 435 440 445

Glu Ala Val Gly Arg His Ile Phe Ala Lys Val Gln Gln Arg Arg Gln 450 455 460

Glu Leu Glu Gln Val Leu Gly Ile Arg Leu Thr 465 470 475

<210> 16

<211> 1235

<212> PRT

<213> Homo sapiens

<400> 16

Met Ala Thr Gly Thr Gly Lys His Lys Leu Leu Ser Thr Gly Pro Thr 1 10 15

Glu Pro Trp Ser Ile Arg Glu Lys Leu Cys Leu Ala Ser Ser Val Met

30 20

Arg Ser Gly Asp Gln Asn Trp Val Ser Val Ser Arg Ala Ile Lys Pro

Phe Ala Glu Pro Gly Arg Pro Pro Asp Trp Phe Ser Gln Lys His Cys

Ala Ser Gln Tyr Ser Glu Leu Leu Glu Thr Thr Glu Thr Pro Lys Arg

Lys Arg Gly Glu Lys Gly Glu Val Val Glu Thr Val Glu Asp Val Ile

Val Arg Lys Leu Thr Ala Glu Arg Val Glu Glu Leu Lys Lys Val Ile 105 100

Lys Glu Thr Gln Glu Arg Tyr Arg Arg Leu Lys Arg Asp Ala Glu Leu

Ile Gln Ala Gly His Met Asp Ser Arg Leu Asp Glu Leu Cys Asn Asp 135

Ile Ala Thr Lys Lys Leu Glu Glu Glu Glu Ala Glu Val Lys Arg

Lys Ala Thr Asp Ala Ala Tyr Gln Ala Arg Gln Ala Val Lys Thr Pro

Pro Arg Arg Leu Pro Thr Val Met Val Arg Ser Pro Ile Asp Ser Ala

Ser Pro Gly Gly Asp Tyr Pro Leu Gly Asp Leu Thr Pro Thr Met 195 200

Glu Glu Ala Thr Ser Gly Val Asn Glu Ser Glu Met Ala Val Ala Ser 210 215

Gly His Leu Asn Ser Thr Gly Val Leu Leu Glu Val Gly Gly Val Leu

Pro Met Ile His Gly Gly Glu Ile Gln Gln Thr Pro Asn Thr Val Ala

Ala Ser Pro Ala Ala Ser Gly Ala Pro Thr Leu Ser Arg Leu Leu Glu 265

Ala Gly Pro Thr Gln Phe Thr Thr Pro Leu Ala Ser Phe Thr Thr Val

Ala Ser Glu Pro Pro Val Lys Leu Val Pro Pro Pro Val Glu Ser Val 300

Ser Gln Ala Thr Ile Val Met Met Pro Ala Leu Pro Ala Pro Ser Ser 320

Ala Pro Ala Val Ser Thr Thr Glu Ser Val Ala Pro Val Ser Gln Pro 325

Asp Asn Cys Val Pro Met Glu Ala Val Gly Asp Pro His Thr Val Thr 340 345 350

Val Ser Met Asp Ser Ser Glu Ile Ser Met Ile Ile Asn Ser Ile Lys 355 360 365

Glu Glu Cys Phe Arg Ser Gly Val Ala Glu Ala Pro Val Gly Ser Lys 370 375 380

Ala Pro Ser Ile Asp Gly Lys Glu Glu Leu Asp Leu Ala Glu Lys Met 385 390 395 400

Asp Ile Ala Val Ser Tyr Thr Gly Glu Glu Leu Asp Phe Glu Thr Val 405 410 415

Gly Asp Ile Ile Ala Ile Ile Glu Asp Lys Val Asp Asp His Pro Glu 420 425 430

Val Leu Asp Val Ala Ala Val Glu Ala Ala Leu Ser Phe Cys Glu Glu 435 440 445

Asn Asp Asp Pro Gln Ser Leu Pro Gly Pro Trp Glu His Pro Ile Gln 450 460

Gln Glu Arg Asp Lys Pro Val Pro Leu Pro Ala Pro Glu Met Thr Val 465 470 475 480

Lys Gln Glu Arg Leu Asp Phe Glu Glu Thr Glu Asn Lys Gly Ile His 485 490 495

Glu Leu Val Asp Ile Arg Glu Pro Ser Ala Glu Ile Lys Val Glu Pro 500 505 510

Ala Glu Pro Glu Pro Val Ile Ser Gly Ala Glu Ile Val Ala Gly Val 515 520 525

Val Pro Ala Thr Ser Met Glu Pro Pro Glu Leu Arg Ser Gln Asp Leu 530 540

Asp Glu Glu Leu Gly Ser Thr Ala Ala Gly Glu Ile Val Glu Ala Asp 545 550 555 560

Val Ala Ile Gly Lys Gly Asp Glu Thr Pro Leu Thr Asn Val Lys Thr 565 570 575

Glu Ala Ser Pro Glu Ser Met Leu Ser Pro Ser His Gly Ser Asn Pro 580 585 590

Ile Glu Asp Pro Leu Glu Ala Glu Thr Gln His Lys Phe Glu Met Ser 595 600 605

Asp Ser Leu Lys Glu Glu Ser Gly Thr Ile Phe Gly Ser Gln Ile Lys 610 615 620

Asp Ala Pro Gly Glu Asp Glu Glu Glu Asp Gly Val Ser Glu Ala Ala 625 630 635 640

Ser Leu Glu Glu Pro Lys Glu Glu Asp Gln Gly Glu Gly Tyr Leu Ser 645 650 655

Glu Met Asp Asn Glu Pro Pro Val Ser Glu Ser Asp Asp Gly Phe Ser 660 665 670

Ile His Asn Ala Thr Leu Gln Ser His Thr Leu Ala Asp Ser Ile Pro 675 680 685

Ser Ser Pro Ala Ser Ser Gln Phe Ser Val Cys Ser Glu Asp Gln Glu 690 695 700

Ala Ile Gln Ala Gln Lys Ile Trp Lys Lys Ala Ile Met Leu Val Trp 705 710 715 720

Arg Ala Ala Asn His Arg Tyr Ala Asn Val Phe Leu Gln Pro Val 725 730 735

Thr Asp Asp Ile Ala Pro Gly Tyr His Ser Ile Val Gln Arg Pro Met 740 745 750

Asp Leu Ser Thr Ile Lys Lys Asn Ile Glu Asn Gly Leu Ile Arg Ser 755 760 765

Thr Ala Glu Phe Gln Arg Asp Ile Met Leu Met Phe Gln Asn Ala Val 770 775 780

Met Tyr Asn Ser Ser Asp His Asp Val Tyr His Met Ala Val Glu Met 785 790 795 800

Gln Arg Asp Val Leu Glu Gln Ile Gln Gln Phe Leu Ala Thr Gln Leu 805 810 815

Ile Met Gln Thr Ser Glu Ser Gly Ile Ser Ala Lys Ser Leu Arg Gly 820 825 830

Arg Asp Ser Thr Arg Lys Gln Asp Ala Ser Glu Lys Asp Ser Val Pro 835 840 845

Met Gly Ser Pro Ala Phe Leu Leu Ser Leu Phe Met Gly His Glu Trp 850 860

Val Trp Leu Asp Ser Glu Gln Asp His Pro Asn Asp Ser Glu Leu Ser 865 870 875 880

Asn Asp Cys Arg Ser Leu Phe Ser Ser Trp Asp Ser Ser Leu Asp Leu 885 890 895

Asp Val Gly Asn Trp Arg Glu Thr Glu Asp Pro Glu Ala Glu Glu Leu 900 905 910

Glu Glu Ser Ser Pro Glu Arg Glu Pro Ser Glu Leu Leu Val Gly Asp 915 920 925

Gly Gly Ser Glu Glu Ser Gln Glu Ala Ala Arg Lys Ala Ser His Gln 930 935 940

Asn Leu Leu His Phe Leu Ser Glu Val Ala Tyr Leu Met Glu Pro Leu 945 950 955 960

Cys Ile Ser Ser Asn Glu Ser Ser Glu Gly Cys Cys Pro Pro Ser Gly 965 970 975

Thr Arg Gln Glu Gly Arg Glu Ile Lys Ala Ser Glu Gly Glu Arg Glu 980 985 990

Leu Cys Arg Glu Thr Glu Glu Leu Ser Ala Lys Gly Asp Pro Leu Val995 1000 1005

Ala Glu Lys Pro Leu Gly Glu Asn Gly Lys Pro Glu Val Ala Ser 1010 1015 1020

Ala Pro Ser Val Ile Cys Thr Val Gln Gly Leu Leu Thr Glu Ser 1025 1030 1035

Glu Glu Gly Glu Ala Gln Gln Glu Ser Lys Gly Glu Asp Gln Gly 1040 1045 1050

Glu Val Tyr Val Ser Glu Met Glu Asp Gln Pro Pro Ser Gly Glu 1055 1060 1065

Cys Asp Asp Ala Phe Asn Ile Lys Glu Thr Pro Leu Val Asp Thr 1070 1075 1080

Leu Phe Ser His Ala Thr Ser Ser Lys Leu Thr Asp Leu Ser Gln 1085 1090 1095

Asp Asp Pro Val Gln Asp His Leu Leu Phe Lys Lys Thr Leu Leu 1100 1105 1110

Pro Val Trp Lys Met Ile Ala Ser His Arg Phe Ser Ser Pro Phe

1115 1120 1125

Leu Lys Pro Val Ser Glu Arg Gln Ala Pro Gly Tyr Lys Asp Val 1130 1135 1140

Val Lys Arg Pro Met Asp Leu Thr Ser Leu Lys Arg Asn Leu Ser 1145 1150 1155

Lys Gly Arg Ile Arg Thr Met Ala Gln Phe Leu Arg Asp Leu Met 1160 1165 1170

Leu Met Phe Gln Asn Ala Val Met Tyr Asn Asp Ser Asp His His 1175 1180 1185

Val Tyr His Met Ala Val Glu Met Arg Arg Glu Val Leu Glu Gln 1190 1200

Ile Glm Val Leu Asn Ile Trp Leu Asp Lys Arg Lys Gly Ser Ser 1205 1210 1215

Ser Leu Glu Gly Glu Pro Ala Asn Pro Val Asp Asp Gly Lys Pro 1220 1225 1230

Val Phe 1235

<210> 17

<211> 513

<212> PRT

<213> Homo sapiens

<400> 17

Met Ala Glu Val Gly Glu Ile Ile Glu Gly Cys Arg Leu Pro Val Leu 1 5 10 15

Arg Arg Asn Gln Asp Asn Glu Asp Glu Trp Pro Leu Ala Glu Ile Leu 20 25 30

Ser Val Lys Asp Ile Ser Gly Arg Lys Leu Phe Tyr Val His Tyr Ile 35 40 45

Asp Phe Asn Lys Arg Leu Asp Glu Trp Val Thr His Glu Arg Leu Asp 50 55 60

Leu Lys Lys Ile Gln Phe Pro Lys Lys Glu Ala Lys Thr Pro Thr Lys 65 70 75 80

Asn Gly Leu Pro Gly Ser Arg Pro Gly Ser Pro Glu Arg Glu Val Pro 85 90 95

Ala Ser Ala Gln Ala Ser Gly Lys Thr Leu Pro Ile Pro Val Gln Ile 100 105 110 Thr Leu Arg Phe Asn Leu Pro Lys Glu Arg Glu Ala Ile Pro Gly Gly 115 120 125

Glu Pro Asp Gln Pro Leu Ser Ser Ser Ser Cys Leu Gln Pro Asn His 130 135 140

Arg Ser Thr Lys Arg Lys Val Glu Val Val Ser Pro Ala Thr Pro Val 145 150 155 160

Pro Ser Glu Thr Ala Pro Ala Ser Val Phe Pro Gln Asn Gly Ala Ala 165 170 175

Arg Arg Ala Val Ala Ala Gln Pro Gly Arg Lys Arg Lys Ser Asn Cys 180 185 190

Leu Gly Thr Asp Glu Asp Ser Gln Asp Ser Ser Asp Gly Ile Pro Ser 195 200 205

Ala Pro Arg Met Thr Gly Ser Leu Val Ser Asp Arg Ser His Asp Asp 210 215 220

Ile Val Thr Arg Met Lys Asn Ile Glu Cys Ile Glu Leu Gly Arg His 225 230 235 240

Arg Leu Lys Pro Trp Tyr Phe Ser Pro Tyr Pro Gln Glu Leu Thr Thr 245 250 255

Leu Pro Val Leu Tyr Leu Cys Glu Phe Cys Leu Lys Tyr Gly Arg Ser 260 265 270

Leu Lys Cys Leu Gln Arg His Leu Thr Lys Cys Asp Leu Arg His Pro 275 280 285

Pro Gly Asn Glu Ile Tyr Arg Lys Gly Thr Ile Ser Phe Phe Glu Ile 290 295 300

Asp Gly Arg Lys Asn Lys Ser Tyr Ser Gln Asn Leu Cys Leu Leu Ala 305 310 315 320

Lys Cys Phe Leu Asp His Lys Thr Leu Tyr Tyr Asp Thr Asp Pro Phe 325 330 335

Leu Phe Tyr Val Met Thr Glu Tyr Asp Cys Lys Gly Phe His Ile Val 340 345 350

Gly Tyr Phe Ser Lys Glu Lys Glu Ser Thr Glu Asp Tyr Asn Val Ala 355 360 365

Cys Ile Leu Thr Leu Pro Pro Tyr Gln Arg Arg Gly Tyr Arg Lys Leu 370 375 380

Leu Ile Glu Phe Ser Tyr Glu Leu Ser Lys Val Glu Gly Lys Thr Gly Thr Pro Glu Lys Pro Leu Ser Asp Leu Gly Leu Leu Ser Tyr Arg Ser Tyr Trp Ser Gln Thr Ile Leu Glu Ile Leu Met Gly Leu Lys Ser Glu Ser Gly Glu Arg Pro Gln Ile Thr Ile Asn Glu Ile Ser Glu Ile Thr Ser Ile Lys Lys Glu Asp Val Ile Ser Thr Leu Gln Tyr Leu Asn Leu Ile Asn Tyr Tyr Lys Gly Gln Tyr Ile Leu Thr Leu Ser Glu Asp Ile 470 475 Val Asp Gly His Glu Arg Ala Met Leu Lys Arg Leu Leu Arg Ile Asp Ser Lys Cys Leu His Phe Thr Pro Lys Asp Trp Ser Lys Arg Gly Lys 505 Trp <210> 18 <211> 364 <212> PRT <213> Homo sapiens <400> 18 Met Ser Leu Ala Gly Gly Arg Ala Pro Arg Lys Thr Ala Gly Asn Arg 10 Leu Ser Gly Leu Leu Glu Ala Glu Glu Glu Asp Glu Phe Tyr Gln Thr Thr Tyr Gly Gly Phe Thr Glu Glu Ser Gly Asp Asp Glu Tyr Gln Gly 35 Asp Gln Ser Asp Thr Glu Asp Glu Val Asp Ser Asp Phe Asp Ile Asp Glu Gly Asp Glu Pro Ser Ser Asp Gly Glu Ala Glu Glu Pro Arg Arg Lys Arg Arg Val Val Thr Lys Ala Tyr Lys Glu Pro Leu Lys Ser Leu Arg Pro Arg Lys Val Asn Thr Pro Ala Gly Ser Ser Gln Lys Ala Arg Glu Glu Lys Ala Leu Leu Pro Leu Glu Leu Gln Asp Asp Gly Ser Asp

Ser Arg Lys Ser Met Arg Gln Ser Thr Ala Glu His Thr Arg Gln Thr 130 135 140

Phe Leu Arg Val Gln Glu Arg Gln Gly Gln Ser Arg Arg Lys Gly 145 150 155 160

Pro His Cys Glu Arg Pro Leu Thr Gln Glu Glu Leu Leu Arg Glu Ala 165 170 175

Lys Ile Thr Glu Glu Leu Asn Leu Arg Ser Leu Glu Thr Tyr Glu Arg 180 185 190

Leu Glu Ala Asp Lys Lys Gln Val His Lys Lys Arg Lys Cys Pro 195 200 205

Gly Pro Ile Ile Thr Tyr His Ser Val Thr Val Pro Leu Val Gly Glu 210 215 220

Pro Gly Pro Lys Glu Glu Asn Val Asp Ile Glu Gly Leu Asp Pro Ala 225 230 235 240

Pro Ser Val Ser Ala Leu Thr Pro His Ala Gly Thr Gly Pro Val Asn 245 250 255

Pro Pro Ala Arg Cys Ser Arg Thr Phe Ile Thr Phe Ser Asp Ala 260 265 270

Thr Phe Glu Glu Trp Phe Pro Gln Gly Arg Pro Pro Lys Val Pro Val 275 280 285

Arg Glu Val Cys Pro Val Thr His Arg Pro Ala Leu Tyr Arg Asp Pro 290 295 300

Val Thr Asp Ile Pro Tyr Ala Thr Ala Arg Ala Phe Lys Ile Ile Arg 305 310 315 320

Glu Ala Tyr Lys Lys Tyr Ile Thr Ala His Gly Leu Pro Pro Thr Ala 325 330 335

Ser Ala Leu Gly Pro Gly Pro Pro Pro Pro Glu Pro Leu Pro Gly Ser 340 345 350

Gly Pro Arg Ala Leu Arg Gln Lys Ile Val Ile Lys 355 360